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With international search report.

(54) Title: 87 HUMAN SECRETED PROTEINS

(57) Abstract

The present invention relates to 87 novel human secreted proteins and isolated nucleic acids containing the coding regions of the genes encoding such proteins. Also provided are vectors, host cells, antibodies, and recombinant methods for producing human secreted proteins. The invention further relates to diagnostic and therapeutic methods useful for diagnosing and treating disorders related to these novel human secreted proteins.

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WO 98/42738 __PCT/US98/05311

87 Human Secreted Proteins

Field of the Invention

This invention relates to newly identified polynucleotides and the polypeptides encoded by these polynucleotides, uses of such polynucleotides and polypeptides, and their production.

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Background of the Invention

Unlike bacterium, which exist as a single compartment surrounded by a membrane, human cells and other eucaryotes are subdivided by membranes into many functionally distinct compartments. Each membrane-bounded compartment, or organelle, contains different proteins essential for the function of the organelle. The cell uses "sorting signals," which are amino acid motifs located within the protein, to target proteins to particular cellular organelles.

One type of sorting signal, called a signal sequence, a signal peptide, or a leader sequence, directs a class of proteins to an organelle called the endoplasmic reticulum (ER). The ER separates the membrane-bounded proteins from all other types of proteins. Once localized to the ER, both groups of proteins can be further directed to another organelle called the Golgi apparatus. Here, the Golgi distributes the proteins to vesicles, including secretory vesicles, the cell membrane, lysosomes, and the other organelles.

Proteins targeted to the ER by a signal sequence can be released into the extracellular space as a secreted protein. For example, vesicles containing secreted proteins can fuse with the cell membrane and release their contents into the extracellular space - a process called exocytosis. Exocytosis can occur constitutively or after receipt of a triggering signal. In the latter case, the proteins are stored in secretory vesicles (or secretory granules) until exocytosis is triggered. Similarly, proteins residing on the cell membrane can also be secreted into the extracellular space by proteolytic cleavage of a "linker" holding the protein to the membrane.

Despite the great progress made in recent years, only a small number of genes encoding human secreted proteins have been identified. These secreted proteins include the commercially valuable human insulin, interferon, Factor VIII, human growth hormone, tissue plasminogen activator, and erythropoeitin. Thus, in light of the pervasive role of secreted proteins in human physiology, a need exists for identifying and characterizing novel human secreted proteins and the genes that encode them. This knowledge will allow one to detect, to treat, and to prevent medical disorders by using secreted proteins or the genes that encode them.

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Summary of the Invention

The present invention relates to novel polynucleotides and the encoded polypeptides. Moreover, the present invention relates to vectors, host cells, antibodies, and recombinant methods for producing the polypeptides and polynucleotides. Also provided are diagnostic methods for detecting disorders related to the polypeptides, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying binding partners of the polypeptides.

Detailed Description

Definitions

The following definitions are provided to facilitate understanding of certain terms used throughout this specification.

In the present invention, "isolated" refers to material removed from its original environment (e.g., the natural environment if it is naturally occurring), and thus is altered "by the hand of man" from its natural state. For example, an isolated polynucleotide could be part of a vector or a composition of matter, or could be contained within a cell, and still be "isolated" because that vector, composition of matter, or particular cell is not the original environment of the polynucleotide.

In the present invention, a "secreted" protein refers to those proteins capable of being directed to the ER, secretory vesicles, or the extracellular space as a result of a signal sequence, as well as those proteins released into the extracellular space without necessarily containing a signal sequence. If the secreted protein is released into the extracellular space, the secreted protein can undergo extracellular processing to produce a "mature" protein. Release into the extracellular space can occur by many mechanisms, including exocytosis and proteolytic cleavage.

As used herein, a "polynucleotide" refers to a molecule having a nucleic acid sequence contained in SEQ ID NO:X or the cDNA contained within the clone deposited with the ATCC. For example, the polynucleotide can contain the nucleotide sequence of the full length cDNA sequence, including the 5' and 3' untranslated sequences, the coding region, with or without the signal sequence, the secreted protein coding region, as well as fragments, epitopes, domains, and variants of the nucleic acid sequence. Moreover, as used herein, a "polypeptide" refers to a molecule having the translated amino acid sequence generated from the polynucleotide as broadly defined.

In the present invention, the full length sequence identified as SEQ ID NO:X was often generated by overlapping sequences contained in multiple clones (contig

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analysis). A representative clone containing all or most of the sequence for SEQ ID NO:X was deposited with the American Type Culture Collection ("ATCC"). As shown in Table 1, each clone is identified by a cDNA Clone ID (Identifier) and the ATCC Deposit Number. The ATCC is located at 10801 University Boulevard, Manassas, Virginia 20110-2209, USA. The ATCC deposit was made pursuant to the terms of the Budapest Treaty on the international recognition of the deposit of microorganisms for purposes of patent procedure.

A "polynucleotide" of the present invention also includes those polynucleotides capable of hybridizing, under stringent hybridization conditions, to sequences contained in SEQ ID NO:X, the complement thereof, or the cDNA within the clone deposited with the ATCC. "Stringent hybridization conditions" refers to an overnight incubation at 42° C in a solution comprising 50% formamide, 5x SSC (750 mM NaCl, 75 mM sodium citrate), 50 mM sodium phosphate (pH 7.6), 5x Denhardt's solution, 10% dextran sulfate, and 20 µg/ml denatured, sheared salmon sperm DNA, followed by washing the filters in 0.1x SSC at about 65°C.

Also contemplated are nucleic acid molecules that hybridize to the polynucleotides of the present invention at lower stringency hybridization conditions. Changes in the stringency of hybridization and signal detection are primarily accomplished through the manipulation of formanide concentration (lower percentages of formanide result in lowered stringency); salt conditions, or temperature. For example, lower stringency conditions include an overnight incubation at 37°C in a solution comprising 6X SSPE (20X SSPE = 3M NaCl; 0.2M NaH₂PO₄; 0.02M EDTA, pH 7.4), 0.5% SDS, 30% formanide, 100 ug/ml salmon sperm blocking DNA; followed by washes at 50°C with 1XSSPE, 0.1% SDS. In addition, to achieve even lower stringency, washes performed following stringent hybridization can be done at higher salt concentrations (e.g. 5X SSC).

Note that variations in the above conditions may be accomplished through the inclusion and/or substitution of alternate blocking reagents used to suppress background in hybridization experiments. Typical blocking reagents include Denhardt's reagent, BLOTTO, heparin, denatured salmon sperm DNA, and commercially available proprietary formulations. The inclusion of specific blocking reagents may require modification of the hybridization conditions described above, due to problems with compatibility.

Of course, a polynucleotide which hybridizes only to polyA+ sequences (such as any 3' terminal polyA+ tract of a cDNA shown in the sequence listing), or to a

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complementary stretch of T (or U) residues, would not be included in the definition of "polynucleotide," since such a polynucleotide would hybridize to any nucleic acid molecule containing a poly (A) stretch or the complement thereof (e.g., practically any double-stranded cDNA clone).

The polynucleotide of the present invention can be composed of any polyribonucleotide or polydeoxribonucleotide, which may be unmodified RNA or DNA or modified RNA or DNA. For example, polynucleotides can be composed of single-and double-stranded DNA, DNA that is a mixture of single- and double-stranded regions, single- and double-stranded RNA, and RNA that is mixture of single- and double-stranded regions, hybrid molecules comprising DNA and RNA that may be single-stranded or, more typically, double-stranded or a mixture of single- and double-stranded regions. In addition, the polynucleotide can be composed of triple-stranded regions comprising RNA or DNA or both RNA and DNA. A polynucleotide may also contain one or more modified bases or DNA or RNA backbones modified for stability or for other reasons. "Modified" bases include, for example, tritylated bases and unusual bases such as inosine. A variety of modifications can be made to DNA and RNA; thus, "polynucleotide" embraces chemically, enzymatically, or metabolically modified forms.

The polypeptide of the present invention can be composed of amino acids joined to each other by peptide bonds or modified peptide bonds, i.e., peptide isosteres, and may contain amino acids other than the 20 gene-encoded amino acids. The polypeptides may be modified by either natural processes, such as posttranslational processing, or by chemical modification techniques which are well known in the art. Such modifications are well described in basic texts and in more detailed monographs, as well as in a voluminous research literature. Modifications can occur anywhere in a polypeptide, including the peptide backbone, the amino acid side-chains and the amino or carboxyl termini. It will be appreciated that the same type of modification may be present in the same or varying degrees at several sites in a given polypeptide. Also, a given polypeptide may contain many types of modifications. Polypeptides may be branched, for example, as a result of ubiquitination, and they may be cyclic, with or without branching. Cyclic, branched, and branched cyclic polypeptides may result from posttranslation natural processes or may be made by synthetic methods. Modifications include acetylation, acylation, ADP-ribosylation, amidation, covalent attachment of flavin, covalent attachment of a heme moiety, covalent attachment of a nucleotide or nucleotide derivative, covalent attachment of a lipid or lipid derivative, covalent attachment of phosphotidylinositol, cross-linking, cyclization, disulfide bond formation, demethylation, formation of covalent cross-links, formation of cysteine,

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formation of pyroglutamate, formylation, gamma-carboxylation, glycosylation, GPI anchor formation, hydroxylation, iodination, methylation, myristoylation, oxidation, pegylation, proteolytic processing, phosphorylation, prenylation, racemization, selenoylation, sulfation, transfer-RNA mediated addition of amino acids to proteins such as arginylation, and ubiquitination. (See, for instance, PROTEINS - STRUCTURE AND MOLECULAR PROPERTIES, 2nd Ed., T. E. Creighton, W. H. Freeman and Company, New York (1993); POSTTRANSLATIONAL COVALENT MODIFICATION OF PROTEINS, B. C. Johnson, Ed., Academic Press, New York, pgs. 1-12 (1983); Seifter et al., Meth Enzymol 182:626-646 (1990); Rattan et al., Ann NY Acad Sci 663:48-62 (1992).)

"SEQ ID NO:X" refers to a polynucleotide sequence while "SEQ ID NO:Y" refers to a polypeptide sequence, both sequences identified by an integer specified in Table 1.

"A polypeptide having biological activity" refers to polypeptides exhibiting activity similar, but not necessarily identical to, an activity of a polypeptide of the present invention, including mature forms, as measured in a particular biological assay, with or without dose dependency. In the case where dose dependency does exist, it need not be identical to that of the polypeptide, but rather substantially similar to the dose-dependence in a given activity as compared to the polypeptide of the present invention (i.e., the candidate polypeptide will exhibit greater activity or not more than about 25-fold less and, preferably, not more than about tenfold less activity, and most preferably, not more than about three-fold less activity relative to the polypeptide of the present invention.)

25 Polynucleotides and Polypeptides of the Invention

FEATURES OF PROTEIN ENCODED BY GENE NO: 1

The translation product of this gene shares sequence homology with nucleolin, which is thought to be important in macromolecule binding, as well as some membrane proteins. Preferred polypeptide fragments comprise the amino acid sequence:

DPEAADSGEPQNKRTPDLPEEEYVKEEIQENEEAVKKMLVEATREFEEVVVDES (SEQ ID NO:239); QKLKRKAEEDPEAADSGEPQNKRTPDLPEEEYVKEEIQENEE AVKKMLVEATREFEEVVVDES (SEQ ID NO:240); KAMEKSSLTQHSWQSLKDR YLKHLRGQEHKYLLGDAPVSPSSQKLKRKAEEDPEAADSGEPQNKRTPDLPEE EYVKEEIQENEEAVKKMLVEATREFEEVVVDESPPDFEIHI (SEQ ID NO:241). Also preferred are the polynucleotide fragments encoding these polypeptide fragments.

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This gene maps to chromosome 16, and therefore can be used as a marker in linkage analysis for chromosome 16.

This gene is expressed primarily in brain and kidney and to a lesser extent in wide range of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cell-cell interaction or cell-matrix interaction. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the brain and kidney, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and kidney, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:125 as residues: Met-1 to Trp-10.

The tissue distribution and homology to nucleolin indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment/diagnosis of diseases involving cell-cell interaction or cell-extracellular matrix interaction.

FEATURES OF PROTEIN ENCODED BY GENE NO: 2

The translation product of this gene shares sequence homology with a porcine zona pellucida protein ZPDS.1711. (See Accession No. R39356.) These two proteins have weak homology with *Drosophila* commissureless and metal homeostasis proteins which are thought to be important in controlling growth cone guidance across the CNS midline and protecting cells against reactive oxygen toxicity. thus, based on homology, it is likely that this gene also be involved in development. Preferred polypeptide fragments comprise the amino acid sequence: LPSYDEAERTKAEATIPLVPGRDEDF VGRDDFDDADQLRIGNDGIFMLTFFMAFLFNWIGFFLSFCLTTSAAGRYGAISG FGLSLIKWILIVRFSTYFPGYFDGQYWLWWVFLVLGFLLFLRGFINYAKVRKM PETFSNLPRTRVLFI (SEQ ID NO:242); and/or AGRYGAISGFGLSLIKWILIVRFS (SEQ ID NO:243). Also preferred are polynucleotide fragments encoding these polypeptide fragments. This gene maps to chromosome 5, and therefore can be used in linkage analysis as a marker for chromosome 5.

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This gene is expressed primarily in kidney, adrenal gland, brain and to a lesser extent in wide range of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, fertilization control or tissue damages by metabolites or other toxic agents. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive and urosecretion system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., kidney, adrenal gland, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to zona pellucida protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for fertility control such as controceptive development. The homology with metal homeostasis and commissureless genes indicates the gene's function in spermatozoa guidance and protection. It would also be useful for the treatment/diagnosis of tissue damages caused by toxic metabolites and other agents since the gene product is also expressed in urosecretive tissues.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 3

This gene is expressed primarily in liver and to a lesser extent in placenta. Preferred polypeptide fragments comprise the amino acid sequence: MKHLSAWNFT KLTFLQLWEI FEGSVENCQTLTSYSKLQIKYTFSRGSTFYI (SEQ ID NO:244). Also preferred are polynucleotide fragments encoding these polypeptide fragments.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, digestive and nutrient transport/utilization disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the digestive and

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circulatory system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., liver, and placenta, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in liver and placenta indicates that the protein product is either an extracellular enzyme or a molecule carrier. Therefore, polynucleotides and polypeptides corresponding to this gene are useful for diagnosis/treatment of digestive and nutrient transport/utilization disorders, including malabsorption and malnutrition.

FEATURES OF PROTEIN ENCODED BY GENE NO: 4

This gene shares homology with the sap47 gene of Drosophila melanogaster, a gene which codes for a conserved neuronal protein associated with synaptic terminals. (See Mol. Brain Res. 32:45-54 (1995); see also, Accession No. 929571.) Thus, based on 15 homology, the gene of the present invention also should be associated with synaptic terminals. Preferred polypeptide fragments comprise the amino acid sequence: FSSDFRTSPWESRRVESKATSARCGLWGSGPRRRPASGMFRGLSSWLGLQQP VAGGGQPNGDAPPEQPSETVAESAEEELQQAGDQELLHQAKDFGNYLFNFASA ATKKITESVAETAQTIKKSVEEGKIDGIIDKTIIGDFQKEQKKFVEEQHTKKSEA 20 AVPPWVDTNDEETIQQQILALSADKRNFLRDPPAGVQFNFDFDQMYPVALVML (SEQ ID NO:245); MRFALVPKLVKEEVFWRNYFYRVSLIKQSAQLTALAAQQQA AGKGGEEQ (SEQ ID NO:246); STSPGVSEFVSDAFDACNLNQEDLRKEMEQL VLDKKQEETAVLEEDSADWEKELQQELQEYEVVTESEKRDENWDK (SEQ ID NO:247); SPWESRRVESKATSARCGLWGSGPRRRPASGMFRGLSSWLGLQQ 25 PVAGGGQPNGDAPPEQPS (SEQ ID NO:248); PVAGGGQPNGDAPPEQPSETV ESAEEELQQAGDQELLHQAKDFGNYLFNFASAATKKITESVAE (SEQ ID NO: 249); and/or FQKEQKKFVEEQHTKKSEAAVPPWVDTNDEETIQQQILALSADKR NFLRDPPAGVQFNFDFDQMYPVALVML (SEQ ID NO:250). Also preferred are polynucleotide fragments encoding these polypeptide fragments. 30

This gene is expressed primarily in kidney pyramids and to a lesser extent in lung and other tissues of various types. This gene fluxes calcium in human aortic smooth muscle cells, and therefore is involved in signal transduction.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, renal and nervous disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the kidney and/or nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., kidney, lung, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in kidney and lung and homology with sap47 indicates that the protein product has regulatory or direct functions in molecular exchange with body fluids and nervous system signaling. Polynucleotides and polypeptides corresponding to this gene are useful for treatment of disorders in kidney and nervous system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 5

The translation product of this gene shares sequence homology with the mouse Ly-9.2 antigen which is thought to be an important cell surface marker in lymphoids, myeloids and hematopoietic progenitors. (See Accession No. gil198932.) Preferred polypeptide fragments comprise the amino acid sequence: PFICVARNPVSRNFSSPI LARKLCEGAA (SEQ ID NO:251); and/or KEDPANTVYSTVEIPKKMENPHSLLT MPDTPRL (SEQ ID NO:252). Also preferred are polynucleotide fragments encoding these polypeptide fragments. Based on homology, it is likely that this gene is also a cell surface marker, involved in hematopoiesis.

This gene is expressed primarily in activated macrophages, monocytes and T-cells and to a lesser extent in spleen and bone marrow.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and hematopoietic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., blood cells, and bone marrow, and cancerous and wounded

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tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:129 as residues: Lys-26 to Tyr-33, Arg-44 to Ile-49, Ser-53 to Lys-71, Lys-86 to Pro-91.

The tissue distribution and homology to Ly-9.2 surface immunoglobulin family indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis of immune and hematopoietic disorders. Polypeptides and polynucleotides corresponding to this gene are also be used as a marker for leukemia or a modulator of the functions of the cells of macrophage/monocyte or T-cell types.

FEATURES OF PROTEIN ENCODED BY GENE NO: 6

The translation product of this gene shares sequence homology with the *Drosophila* glutactin gene which is thought to be important in cell-cell interaction or cell-extracellular matrix contact.

This gene is expressed primarily in colon tissue, aorta endothelial cells and to a lesser extent in skin, breast tissue and T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of these tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the gastrointestinal tract, vascular system or T-cell development. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the digestive system, cardiovascular system, and immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., colon, cardiovascular tissue, skin, mammary tissue, and blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to glutactin indicates that polynucleotides and polypeptides corresponding to this gene are useful for the development and maintenance of the integrity of the basal membrane in the gastrointestinal tract and

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cardiovascular system. The expression in T-cells also indicate the protein may be involved in T-cell adhesion, cell-cell interaction and development.

FEATURES OF PROTEIN ENCODED BY GENE NO: 7

The translation product of this gene shares sequence homology with MURF4 protein, an ATPase homolog, which is thought to be important in ATP hydrolysis.

This gene is expressed primarily in breast tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, breast cancer and non-neoplastic breast diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the breast tissue, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to MURF4 gene indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of neoplastic or non-neoplastic breast diseases because ATPase like protein may be involved in changed metabolic states of the breast.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 8

This gene shares homology to the alcohol dehydrogenase gene. Preferred polypeptide fragments comprise comprise the amino acid sequence: ASAVLLDLPNSG GEAQAKKLGNNCVFAPADVTSEKDVQTALALAKGKFGRVDVAVNCAGIAVAS KTYNLKKGQTHTLEDFQRVLDVNLMGTFNVIRLVAGEMGQNEPDQGGQRGVI INTASVAAFEGQVGQAAYSASKGGIVGMTLPIARDLAPIGIRVMTIAPGLFGTPL LTSLPEKVCNFLASQVPFPSRLGDPAEYAHLVQAIIENPFLNGEVIRLDGAIRMQ P (SEQ ID NO:253); and/or SVAAFEGQVGQAAYSASKGGIVGMTLPIA (SEQ ID NO:254). Polynucleotides encoding these fragements are also encompassed by the invention. Other groups have also recently cloned this gene, recognizing its homology to alcohol dehydrogenase. (See Accession No. 1778355.) Moreover, a second group

recently cloned the mouse homologue of this gene. (See Accession No. 2078284.) They found that the mouse homologue binds to amyloid beta-peptide and mediates neurotoxicity in Alzheimer's disease, calling the protein ERAB. This gene maps to chromosome X, and therefore can be used in linkage analysis as a marker for chromosome X. Therefore, mutations in the translated product of this gene may be involved in Alzheimer's disease in humans, as well as other sex linked diseases. This gene can be used as a diagnostic marker for these diseases.

Preferred epitopes include those comprising a sequence shown in SEQ ID NO:132 as residues: Arg-45 to Ser-53.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 9

The translation product of this gene shares week sequence homology with rat N-methyl-D-aspartate receptor subunit and other proline-rich proteins which are thought to be important in neurotransmission or protein-protein intereaction.

This gene is expressed primarily in synovial hypoxia and to a lesser extent in ovary, senescent cells and brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, synovial hypoxia. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the synovia and brain, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., synovial tissue, ovary and other reproductive tissue, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in synovial hypoxia and nerve tissues, and homology to N-methyl-D-aspartate receptor subunit and other proline-rich proteins indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and intervention of synovial hypoxia and other synovial disorders.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 10

This gene is expressed primarily in prostate and to a lesser extent in placenta and ovary.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, male and female infertility, cancer, and other hyperproliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive system and neoplasia, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., prostate, placenta, ovary and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:134 as residues: Pro-17 to Met-23, Ala-30 to Trp-38, Ile-49 to Trp-54, Lys-68 to Gly-74, Thr-93 to Gly-99, Met-126 to Glu-132, Gly-173 to Ser-178, Lys-205 to Tyr-214.

The tissue distribution of this gene in the prostate, placenta and ovary indicates that this gene product is useful for treatment/diagnosis of male or female infertility, endocrine disorders, fetal deficiencies, ovarian failure, amenorrhea, ovarian cancer, benign prostate hyperplasia, prostate cancer, and other forms of cancer of the reproductive system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 11

This gene is expressed primarily in the thyroid and to a lesser extent in the pineal gland. This gene maps to chromosome 10, and therefore can be used as a marker in linkage analysis for chromosome 10. Preferred polypeptide fragments comprise the amino acid sequence: HPIEWAINAATLSQFY (SEQ ID NO:256); CWIKYCLTLMQN AQLSMQDNIG (SEQ ID NO:257); KVSYLRPLDFEEARELFLLGQHYVF (SEQ ID NO:258); MERRCKMHKRXIAMLEPLTVDLNPQ (SEQ ID NO:259); and/or SHIV KKINNLNKSALKY YQLFLD (SEQ ID NO:260). Also preferred are polynucleotides encoding these polypeptide fragments.

Therefore, polynucleotides and polypeptides of the invention are useful as

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reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune, thyroid and pineal gland disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., thyroid and pineal gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:135 as residues: Ser-2 to Ser-8, Thr-38 to Arg-44.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treating/detecting immune disorders such as arthritis, asthma, immune deficiency diseases (e.g., AIDS), and leukemia, as well as treating/detecting thymus disorders (e.g., Graves Disease, lymphocytic thyroiditis, hyperthyroidism, and hypothyroidism), and treating/detecting pineal gland disorders (e.g., circadian rhythm disturbances associated with shift work, jet lag, blindness, insomnia and old age).

FEATURES OF PROTEIN ENCODED BY GENE NO: 12

This gene is expressed primarily in lung and tonsils.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, pulmonary or immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the pulmonary and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pulmonary tissue, and tonsils, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily

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fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:136 as residues: Glu-28 to Gly-49.

The tissue distribution of this gene only in lung indicates that it could play a role in the treatment/detection of lung lymphoma or sarcoma formation, pulmonary edema and embolism, bronchitis and cystic fibrosis. Its expression in tonsils indicates a potential role in the treatment/detection of immune disorders such as arthritis, asthma, immune deficiency diseases (e.g., AIDS), and leukemia, in addition to the treatment/detection of tonsillitis.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 13

This gene is expressed primarily in lymphoid, myeloid and erythroid cells. Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, hematopoietic and immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the hematopoietic and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, myeloid cells, and bone marrow, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The predominant tissue distribution of this gene in hematopoietic cell types indicates that the gene could be important for the treatment or detection of immune or hematopoietic disorders including arthritis, asthma, immunodeficiency diseases and leukemia. Preferred embodiments of the present invention are polypeptide fragments comprising the amino acid sequence: FTHLSTCLLSLLLVRMSGFLLLARASPSI CALDSSCFVEYCSSYSSSCFLHQHFPSLLDHLCQ (SEQ ID NO:261); or FLLL ARASPSICALDSSCFVQEY (SEQ ID NO:262). Also preferred are polynucleotide fragments encoding these polypeptide fragments.

FEATURES OF PROTEIN ENCODED BY GENE NO: 14

This gene is homologous to the *Drosophila Regena* (Rga) gene. (See Accession No. 1658504.) This *Drosophila* gene is thought to be a homolog of the global negative

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transcriptional regulator NOT2 (CDC36) from yeast, which modifies gene expression and suppresses position effect variegation. Preferred polypeptide fragments comprise the amino acid sequence: PDGRVTNIPQGMVTDQFGMIGLLTFIRAAETDPGMVHL ALGSDLTTLGLNLNS (SEQ ID NO:263); VHLALGSDLTTLGLNLNSPENLYP (SEQ ID NO:265); EDLLFYLYYMNGGDVLQLLAAVELFNRDWRYHKEERVWI TR (SEQ ID NO:264); and/or HNEDFPALPGS (SEQ ID NO:266).

This gene is expressed primarily in placenta and to a lesser extent in infant brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurodegenerative and developmental disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the neurological system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:138 as residues: Leu-9 to Tyr-15, Asp-34 to Gln-46, Pro-51 to Asp-57, Gly-88 to Thr-104, Thr-123 to Ser-128.

The tissue distribution of this gene indicates that it could be used in the detection and/or treatment of neurological disorders such as such as Alzheimer's Disease, Parkinson's Disease, Huntington's Disease, schizophrenia, mania, dementia, paranoia, obsessive compulsive disorder, and panic disorder.

FEATURES OF PROTEIN ENCODED BY GENE NO: 15

This gene is expressed primarily in adrenal gland tumor and osteoclastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, endocrine and bone disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for

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differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine system and in bone, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adrenal gland, and bone, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:139 as residues: Ile-52 to Trp-57.

The tissue distribution of this gene indicates that it may be involved in the treatment and/or detection of adrenal gland tumors, osteosarcomas, endo time disorders and bone disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 16

The translation product of this gene shares sequence homology th the FK506 binding protein, a protein which plays an important role in im success pression. (See Accession No. M75099.) Specifically, a 12-kDa FK506-binding and con (FKBP-12) is a cytosolic receptor for the immunosuppressants FK506 and apamy to (See, Proc. Natl. Acad. Sci. 88: 6677-6681 (1991).) Thus, based on hour every likely that this gene also has immunosuppression activity. Preferred pullyperally rise the amino acid sequence: GRIIDTSLTRDPLVIELGQKQVIPGLIBQS KRRAIIPSH LAYGKRGFPPSVPADAVVQYDVELIALIR (SEQ ID) NO:25 HYTGSLV DGR IIDTS (SEQ ID NO:268). Also preferred are the pois next unents encoding these polypeptides.

This gene is expressed primarily in melanocytess.

Therefore, polynucleotides and polypeptides offthe inlas reagents for differential identification of the tissue(s) of the biological sample and for diagnosis of diseases and condibut are not limited to, cancer and other hyperproliferative disord ptides and antibodies directed to these polypeptides are useful. gical probes for differential identification of the tissue(s) or con-ਾ of disorders of the above tissues or cells, particularly of the ancer, expression of this gene at significantly higher or lower le letected in certain tissues and cell types (e.g., melanocytes, and c. . . . tissues) or bodily fluids (e.g., serum, plasma, urine, synid) or another tissue or cell sample taken from an individual ha lative to

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the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:140 as residues: Ala-118 to Phe-124, Arg-178 to Lys-201.

The tissue distribution and homology to the FK506 binding proteins which are believed to a role in immunosupression mediated by the immunosupressant drugs rapamycin and cyclosporin, indicates that this gene could serve as a novel target for the identification of novel immunosupressant drugs.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 17

The translation product of this gene shares sequence homology with the rat calcium-activated potassium channel rSK3, which is thought to be important in regulating vascular tone. (See Accession No. gil2564072, gil1575663, and gil1575661.) Although homologous to these proteins, this gene contains an 18 amino acid insert, not previously identified in the homologs. Preferred polypeptide fragments comprise the amino acid sequence: CESPESPAQPSGSSLPAWYH (SEQ ID NO:269). Also preferred are the polynucleotide fragments encoding these polypeptides.

This gene is expressed primarily in B-cells, frontal cortex and endothelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cardiovascular (hyper/hypotension, asthma, pulmonary edema, pneumonia, heart disease, restenosis, atherosclerosis, stoke, angina and thrombosis) and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular and nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, brain and other tissue of the nervous system, and endothelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:141 as residues: Glu-72 to Gly-82, His-90 to Val-95, Gln-168 to Lys-174, Val-202 to Ser-212.

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The tissue distribution and homology to calcium-activated potassium channels indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of vascular disorders (hyper/hypotension, athesma, pulmonary edema, pneumonia, heart disease, restenosis, atherosclerosis, stoke, angina and thrombosis).

FEATURES OF PROTEIN ENCODED BY GENE NO: 18

This gene is expressed primarily in smooth muscle and to a lesser extent in brain (amygdala, corpus colosum, hippocampus).

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cardiovascular (hypertension, heart disease, athesma, pulmonary edema, restenosis, atherosclerosis, stoke, angina, thrombosis, and wound healing), and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular and neurological systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., smooth muscle, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:142 as residues: Lys-43 to Arg-49, Tyr-58 to Glu-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of cadiovascular disorders (hypertension, heart disease, athesma, pulmonary edema, restenosis, atherosclerosis, stoke, angina, thrombosis, and wound healing). Expression in brain indicates a role in the treatment and diagnosis of behavioral or neurological disorders, such as depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior.

35 FEATURES OF PROTEIN ENCODED BY GENE NO: 19

This gene is expressed primarily in T-cells (Jurkats, resting, activated, and

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anergic T-cells), endothelial cells, pineal gland, and to a lesser extent in a variety of other tissues and cell types. Preferred polypeptide fragments comprise the amino acid sequence: EEAGAGRRCSHGGARPAGLGNEGLGLGGDPDHTDTGSRSKQRINN WKESKHKVIMASASARGNQDKDAHFPPPSKQSLLFCPKSKLHIHRAEISK (SEQ ID NO:270); and/or SKQRINNWKESKHKVIMASASAR (SEQ ID NO:271). Also preferred are the polynucleotide fragments encoding these polypepides.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation, immune and cardiovascular disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neurological and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, endothelial cells, and pineal gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:143 as residues: Phe-71 to Arg-76, Pro-82 to His-87, Glu-103 to Ala-111.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. In addition, expression in the pineal gland might suggest a role in the diagnosis of specific brain tumors and treatment of neurological disorders. Endothelial cell expression might suggest a role in cadiovascular or respiratory/pulmonary disorders or infections (athesma, pulmonary edema, pneumonia).

FEATURES OF PROTEIN ENCODED BY GENE NO: 20

This gene is expressed primarily in brain and embryo and to a lesser extent in leukocytes. This gene maps to chromosome 15, and therefore can be used as a marker in linkage analysis to chromosome 15.

Therefore, polynucleotides and polypeptides of the invention are useful as

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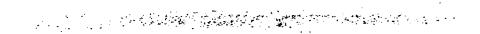
reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, developmental and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO: 144 as residues: Met-1 to Gly-8.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain -- and in particular the fetal brain -- would suggest a possible role in the treatment and diagnosis of developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior).

FEATURES OF PROTEIN ENCODED BY GENE NO: 21

This gene is expressed primarily in brain, kidney, lung, liver, spleen, and a variety of leukocytes (especially T-cells) and to a lesser extent in a variety of other tissues and cell types.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include but are not limited to, leukemias, lymphomas, autoimmune, immunosuppressive, and immunodeficiencies, hematopoietic disorders, as well as renal disorders, and neoplasms. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, partic darly of the renal, pulmonary, immune, and central nervous systems, expression of the significantly higher or lower levels may be routinely detected in certain tissue (e.g.,



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brain and other tissue of the nervous system, kidney, pulmonary tissue, liver, spleen, and blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of renal conditions, such as acture renal failure, kidney fibrosis, and kidney tubule regeneration. The expression in leukocytes and other immune tissues indicates a role in immune disorders including: leukemias. lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain -- and in particular the fetal brain -- indicates a possible role in the treatment and diagnosis of developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior).

FEATURES OF PROTEIN ENCODED BY GENE NO: 22

This gene is expressed primarily in skin (fetal epithelium, keratinocytes and skin). This gene also maps to chromosome 19, and therefore can be used in linkage analysis as a marker for chromosome 19.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, skin cancers (e.g., melanomas), eczema, psoriasis or other disorders of the skin. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the skin, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., skin and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:146 as residues: Pro-28 to Glu-35, Ser-39 to Phe-44, Ala-94 to Gln-99.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of skin cancers (e.g., melanomas), eczema, psoriasis or other disorders of the skin.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 23

This gene maps to chromosome 11. Another group recently isolated this same gene, associating the sequence to the region thought to harbor the gene involved in Multiple Endocrine Neoplasia Type 1, or MEN 1. (See Accession No. 2529721 and Genome Res. 7(7), 725-735 (1997), incorporated herein by reference in its entirety.) Preferred polypeptide fragments comprise the amino acid sequence: LFHWACLNERA AQLPRNTAXAGYQCPSCNGPS (SEQ ID NO:272).

This gene is expressed primarily in epididymus, pineal gland, T-cells, as well as fetal epithelium, lung and kidney.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune, metabolic mediated disorders, and MEN. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, renal, neurological and pulmonary systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., epididymus and other reproductive tissue, pineal gland, T-cells and other blood cells, epithelium, lung, and kidney, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of developmental deficiencies or abnormalities as well as a host of different disorders which arise as a result of conditions in the indicated tissues or cell types. An area of particular interest is in the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain, and in particular the fetal brain, would suggest a possible role in the treatment and diagnosis of

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developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior). Respiratory/pulmonary disorders, such as athesma, pulmonary edema are also potential therapeutic areas, as well as renal conditions such as acute renal failure, kidney fibrosis and kidney tubule regeneration. Moreover, this gene can be used in the treatment and/or detection of MEN I.

FEATURES OF PROTEIN ENCODED BY GENE NO: 24

This gene is expressed primarily in fetal spleen.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, leukemia, lymphoma, AIDS, hematopoeitic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., spleen and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 25

A closely related homolog of this gene was recently cloned by another group, calling the gene CDO, an oncogene-, serum-, and anchorage-regulated member of the Ig/fibronectin type III repeat family. (See Accession No. 2406628, and J. Cell Biol. 138(1): 203-213 (1997), herein incorporated by reference in its entirety.) Preferred polypeptide fragments comprise the amino acid sequence: FYIYYRPTDSDNDSDYKK DMVEGDKYWHSISHLQPETSYDIKMQCFNEGGESEFSNVMICETKARKSSGQP GRLPPPTLAPPQPPLPETIERPVGTGAMVARSSDLPYLIVGVVLGSIVLIIVTFIPF CLWRAWSKQKHTTDLGFPRSALPPSCPYTMVPLGGLPGHQAVDSPTSVASVD

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GPVLM (SEQ ID NO:273); or YIYYRPTDSDNDSDYKKDMVEGDKYWHSISHLQ PETSYDIKMQCFNEGGESEFSNVMICETKARKS (SEQ ID NO:274).

This gene is expressed primarily in fetal lung and kidney, human embryo and osteoclastoma stromal cells and to a lesser extent in a variety of other tissues and cell types.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, developmental disorders and cancers, as well as pulmonary and renal disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the respiratory/pulmonary, skeletal and renal systems, expression of this generat significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., lung, kidney, embryonic tissue, and bone cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder. relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:149 as residues: Thr-5 to Pro-18, Ala-76 to Thr-84.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the detection and treatment of: osteoperosis. fracture, osteosarcoma, ossification, and osteonecrosis, as well as respiratory/pulmonary disorders, such as athesma, pulmonary edema, and renal conditions such as acute renal failure, kidney fibrosis and kidney tubule regeneration.

FEATURES OF PROTEIN ENCODED BY GENE NO: 26

This gene is homologous to the HIV envelope glycoprotein. (See Accession No. 2641463.) Preferred polypeptide fragments comprise the amino acid sequence: NVRALLHRMPEPPKINTAKFNNNKRKNLSL (SEQ ID NO:275).

This gene is expressed primarily in pineal gland and skin, and to a lesser extent in lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

not limited to, neurological and behavior disorders; respiratory/pulmonary disorders, such as athesma, pulmonary edema; skin conditions such as eczema, psoriasis, acne and skin cancer, as well as AIDS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous and respiratory systems, as well as skin and AIDS, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, pineal gland, epidermis, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:150 as residues: Gln-15 to Gln-20.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of conditions which affect the above tissues, such as: skin cancer, eczema, psoriasis, acne, athesma, pulmonary edema, neuro-degenerative or developmental disorders such as Alzheimer's, depression, schizophrenia, dementia, and AIDS.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 27

Preferred polypeptide encoded by this gene comprise the following amino acid sequence: NTNQREALQYAKNFQPFALNHQKDIQVLMGSLVYLRQGIENSPYVHL LDANQWADICDIFTRDACALLGLSVESPLSVSFSAGCVALPALINIKAVIEQRQC TGVWNQKDELPIEVDLGKKCWYHSIFACPILRQQTTDNNPPMKLVCGHIISRD ALNKMFNGSKLKCPYCPMEQSPGDAKQIFF (SEQ ID NO:276). Polynucleotides encoding such polypeptides are also provided as are complementary polynucleotides thereto.

This gene is expressed primarily in liver (adult and fetal) and spleen tissue, and to a lesser extent in placenta, T helper cells, kidney tumor, ovarian tumor, melanocytes and fetal heart.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and developmental diseases and disorders and liver diseases such as liver cancer. Similarly, polypeptides and antibodies directed to these

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polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, circulatory and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., liver, spleen, placenta, blood cells, kidney, ovary and other reproductive tissue, melanocytes, and heart, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of growth, hematopoietic and immune system disorders particularly related to the liver.

15 FEATURES OF PROTEIN ENCODED BY GENE NO: 28

The translation product of this gene shares sequence homology with prostaglandin transporter which is thought to be important in metabolic and endocrine disorders. See, for example, Gastroenterology Oct:109(4):1274-1282 (1995). Preferred polypeptides encoded by this gene comprise the following amino acid sequence: SYLSACFAGCNSTNLTGCACLTTVPAENATVVPGKCPSPGCQEAFLTFLCVMCI CSLIGAMARHP (SEQ ID NO:277); and/or PSVIILIRTVSPGLKSYALGVLFLLLRL LGFIPPPLIFGAGIDSTCLFWSTFCGEQGACVLYDNVV) YLYVSIAIALKSFAFI (SEQ ID NO:278).

This gene is expressed primarily in hematopoietic ans	rain tissues.	
Therefore, polynucleotides and polypeptides of the in-	mion are usef	: as
reagents for differential identification of the tissue(s) or cell) present .	
biological sample and for diagnosis of diseases and condition	hich inclu	ut are
not limited to, metabolic, immune and endocrine diseases and	.078 S	rly.
polypeptides and antibodies directed to these polypeptides are	. m pr	·g
immunological probes for differential identification of the tis	or well.). For a
number of disorders of the above tissues or cells, particularly	di.	mmune
and endocrine systems, expression of this gene at significant		evels
may be routinely detected in certain tissues (e.g., endocrime to		•
tissue, and brain and other tissue of the nervous system, and	•	nded
tissues) or bodily fluids (e.g., serum, plasma, urine, synewis		d) or
another tissue or cell sample taken from an individual haviin		tive to

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the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to prostaglandin (and anion) transporter indicates that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis and treatment of endocrine, metabolic, immune and kidney disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 29

This gene is expressed primarily in early stage human lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, growth and respiratory disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the developmental and respiratory systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:153 as residues: Val-50 to Trp-55.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of respiratory and growth diseases and disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 30

The translation product of this gene shares sequence homology with human DNA helicase which is thought to be important in accurate and complete DNA replication in creation of new cells. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: QSLFTRFVRVGVPTVDLDAQGRARA SLCXXYNWRYKNLGNLPHVQLLPEFSTANAGLLYDFQLINVEDFQGVGESEPN PYFYQNLGEAEYVVALFMYMCLLGYPADKISILTTYNGQKHLIRDIINRRCGNN PLIGRPNKVTTVDRFQGQQNDYILLSLVRTRAVGHLRDVRRLVVAMSRAR (SEQ ID NO:279); and/or LVKEAKIIAMTCTHAALKRHDLVKLGFKYDNILMEE AAQILEIETFIPLLLQNPQDGFSRLKRWIMIGDHHQLPPVI (SEQ ID NO:280).

This gene is expressed primarily in testes tumor and to a lesser extent in adrenal

gland tumor and placenta.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancers and endocrine/growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine, developmental, and reproductive systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., testes and other reproductive tissue, adrenal gland, and placenta, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to DNA helicase indicates that the protein products of this gene are useful for study, treatment, and diagnosis of many cancer types, including testicular cancer; as well as disorders involving endocrine function and normal growth and development.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 31

The translation product of this gene shares sequence homology with BID-apoptotic death gene (mouse), Genbank accession no. PID g1669514, which is thought to be important in programmed cell death.

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This gene is expressed primarily in jurkat membrane bound polysomes and activated neutrophils and to a lesser extent in endothelial cells and human cerebellum.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancers and other proliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, endothelium, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma,

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urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:155 as residues: Glu-4 to Leu-11, Cys-28 to Arg-35, Gln-50 to His-66, Glu-73 to Gln-79, Gly-94 to Ser-100, Arg-114 to Asp-126, Pro-139 to Lys-146.

The tissue distribution and homology to BID-apoptotic death gene indicates that the protein products of this gene are useful for study of cell death, and treatment and diagnosis of proliferative disorders and cancers. Apoptosis - programmed cell death - is a physiological mechanism involved in the deletion of peripheral T lymphocytes of the immune system, and its dysregulation can lead to a number of different pathogenic processes. Diseases associated with increased cell survival, or the inhibition of apoptosis, include cancers (such as follicular lymphomas, carcinomas with p53 mutations, and hormone-dependent tumors, such as breast cancer, prostrate cancer, Kaposiís sarcoma and ovarian cancer); autoimmune disorders (such as systemic lupus erythematosus and immune-related glomerulonephritis rheumatoid arthritis) and viral infections (such as herpes viruses, pox viruses and adenoviruses), inflammation; graft vs. host disease, acute graft rejection, and chronic graft rejection. Diseases associated with increased apoptosis include AIDS; neurodegenerative disorders (such as Alzheimer's disease, Parkinson's disease, Amyotrophic lateral sclerosis, Retinitis pigmentosa, Cerebellar degeneration); myelodysplastic syndromes (such as aplastic anemia), ischemic injury (such as that caused by myocardial infarction, stroke and reperfusion injury), toxin-induced liver disease (such as that caused by alcohol), septic shock, cachexia and anorexia. Thus, the invention provides a method of enhancing apoptosis in an individual by treating the individual with a polypeptide encoded by this gene.

FEATURES OF PROTEIN ENCODED BY GENE NO: 32

The translation product of this gene shares sequence homology with human fructose transporter which is thought to be important in normal metabolic function and activity.

This gene is expressed primarily in T-cell lymphoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, leukemia and other cancers, and metabolic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the hematopoietic, lymph and metabolic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:156 as residues: Pro-22 to Gly-48, Ser-54 to Pro-61.

The tissue distribution indicates that the protein products of this gene are useful for study of mechanisms leading to cancer, treatment and diagnosis of cancerous and pre-cancerous conditions; as well as the study and treatment of various metabolic diseases and disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 33

This gene is expressed primarily in human meningima.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and other disorders of the CNS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the CNS and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., meningima and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:157 as residues: Asn-23 to Pro-31.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of disorders of the CNS and inflammatory responses.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 34

This gene is expressed primarily in activated monocytes and wound healing tissues and to a lesser extent in fetal epithelium.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and inflammatory disorders and wound healing and tissue repair dysfunctions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, epithelial and gastrointestinal systems, and healing wounds, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., monocytes and other blood cells, and epithelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:158 as residues: Ala-28 to Ala-33, Gly-35 to Glu-45.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis, study and treatment of immune and inflammatory disorders and wound healing dysfunctions.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 35

This gene is expressed primarily in human osteosarcoma and prostate cancer.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, skeletal and neoplastic conditions such as bone and prostate cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and skeletal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and prostate, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial

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fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:159 as residues: Ser-14 to Gly-22, Leu-37 to Gln-43.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis and treatment of skeletal disorders and cancer.

FEATURES OF PROTEIN ENCODED BY GENE NO: 36

This gene encodes a protein which is highly homologous to a protein called congenital heart disease protein 5, presumably implicated in congenital heart disease (see Genbank PID g2810996).

This gene is expressed primarily in Hodgkin's lymphoma, erythroleukemia cells, and TNF activated synovial fibroblasts, to a lesser extent in ovarian cancer, cerebellum, spleen, fetal liver and placenta and finally to a lesser extent in various other mesenchymal tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer, immune, hematopoietic and cardiovascular disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, hematopoietic and cardiovascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., heart and other cardiovascular tissue, lymphoid tissue, blood cells, bone marrow, ovary and other reproductive tissue, brain and other tissue of the nervous system, spleen, liver, and mesenchymal tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:160 as residues: Lys-41 to Met-49, Gln-54 to Glu-59, Glu-76 to Thr-88.

The homology of this gene and translation product to congenital heart disease protein 5 indicates a role for this protein in the diagnosis, prognosis and/or treatment of

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heart disease or other cardiovascular related disorders. In addition, predominant expression in cells associated with the immune and hematopoetic system indicates a role for this protein in the treatment, diagnosis and/or prognosis of immune and autoimmune diseases, such as lupus, transplant rejection, allergic reactions, arthritis, asthma, immunodeficiency diseases, leukemia, AIDS, thymus disorders such as Graves Disease, lymphocytic thyroiditis, hyperthyroidism and hypothyroidism, graft versus host reaction, graft versus host disease, transplant rejection, myelogenous leukemia, bone marrow fibrosis, and myeloproliferative disease. The protein could also be used to enhance or protect proliferation, differentiation and functional activation of hematopoietic progenitor cells such as bone marrow cells, which could be useful for cancer patients undergoing chemotherapy or patients undergoing bone marrow transplantation. The protein may also be useful to increase the proliferation of peripheral blood leukocytes, which could be useful in the combat of a range of hematopoietic disorders including immunodeficiency diseases, leukemia, and septicemia.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 37

This gene is expressed primarily in ovarian cancer.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, urogenital neoplasias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., ovary and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:161 as residues: Asn-22 to Asn-27.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis and treatment of ovarian and other tumors.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 38

The translation product of this gene shares sequence homology with zinc finger proteins.

This gene is expressed primarily in various fetal, cancer, and endothelial lines.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., fetal tissue, and endothelial cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of immune and developmental conditions and cancer.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 39

This gene is expressed primarily in fetal, infant, and adult brain and to a lesser extent in other brain and endocrine organs and blastomas.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, brain tumors and neurodegenerative conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, endocrine tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the

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disorder.

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The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of brain cancer and other neurological disorders.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 40

The translation product of this gene shares sequence homology with vesicular glycoproteins and lectins. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: DTYPNEEKQQERVFPXXSAMVNNGSLSYDHER DGRPTELGGCXAIVRNLHYDTFLVIRYVKRHLTIMMDIDGKHEWRDCIEVPGV RLPRGYYFGTSSITGDLSDNHDVISLKLFELTVERTPEEE (SEQ ID NO:281); and/or LKREHSLSKPYQGVGTGSSSLWNLMGNAMVMTQYIRLTPDMQSKQGA LWNRVPCFLRDWELQVHFKIHGQGKKNLHGDGLAIWYT (SEQ ID NO:282).

This gene is expressed primarily in infant brain and to a lesser extent in various normal and transformed neural, endocrine, and immune organs.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurological and neurodevelopmental conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous and hormonal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, endocrine tissue, and tissue and cells of the immune system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:164 as residues: Pro-64 to Gly-71, Gly-94 to Leu-100, Thr-110 to Pro-116, Thr-135 to Arg-145, Glu-164 to Glu-171, Asp-204 to Asp-211, Arg-253 to His-261, Asn-312 to Tyr-323.

The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of mental retardation and other neurological disorders and neoplasias.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 41

This gene displays homology to the glycosyltransferase family, which catalyze the addition of sialic acids to carbohydrate groups which are present on glycoproteins.

This gene is expressed primarily in smooth muscle and to a lesser extent in pineal gland, fetal liver, and infant brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, gastrointestinal injury, inflammatory and neurodegenerative conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., smooth muscle, pineal gland, liver, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:165 as residues: Ser-12 to Trp-21, Arg-24 to Pro-32, Asp-73 to Lys-82, Lys-90 to Ala-97.

The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of neurodegenerative and growth disorders and gastrointestinal repair.

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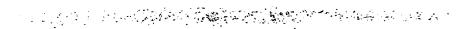
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FEATURES OF PROTEIN ENCODED BY GENE NO: 42

The translation product of this gene shares sequence similarity with metallothionein polypeptides. See, for example, Proc. Natl. Acad. Ser. U.S. A 1992 Jul 15:89(14):6333-6337. Metallothioneins are believed to inhibit neuronal scarvival among other biological functions. Based on the sequence similarity (espectable the conserved cysteine motifs characteristic of the metallothionein family) the trackle on product of this gene is expected to share certain biological activities with other metallothionein polypeptide family. Preferred polypeptides encoded the metallothionein polypeptide family. Preferred polypeptides encoded the second comprise the following amino acid sequence: PGTLQCSALHER ACANC CRD CSPPACQC (SEQ ID NO:283).

This gene is expressed exclusively in placenta and fetal:



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Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, hematopoietic and immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, liver, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to metallothionien indicates that the protein products of this gene are useful for diagnosis and treatment of immune and hematopoietic system disorders and neurological diseases, especially in fetal development.

20 FEATURES OF PROTEIN ENCODED BY GENE NO: 43

Preferred polypeptides encoded by this gene comprise the following amino acid sequence: FLYDVLMXHEAVMRTHQIQLPDPEFPS (SEQ ID NO:284).

This gene is expressed primarily in T-cells and synovial tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune system disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., synovial tissue, and T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

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The tissue distribution indicates that the protein products of this gene are useful for treatment and diagnosis of disorders of the immune system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 44

The translation product of this gene shares sequence similarity with several methyltransferases (e.g., see Genbank gil1065505).

This gene is expressed primarily in ovary, thymus, infant adrenal gland, tissues of the nervous system and the hematopoietic tissue, and to a lesser extent in adipose tissue and many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, disorders of the reproductive system, the endocrine system, the hematopoietic system and the CNS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, endocrine, CNS and reproductive system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., ovary and other reproductive tissue, thymus, adrenal gland, brain and other tissue of the nervous system, hematopoietic tissue, and adipose tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:168 as residues: Ser-3 to Gly-12, Asp-19 to Arg-31, Tyr-70 to Tyr-77, Asn-130 to Lys-140, Pro-165 to Gln-170, Pro-192 to Lys-199, Leu-216 to Glu-227, Glu-254 to Phe-281.

The tissue distribution and homology to methyltransferase indicates that the protein products of this gene are useful for diagnosis and treatment of disorders of the CNS, the hematopoietic system and reproductive organs and tissues. For example, the abundant expression in the ovary may indicate that the gene product can be used as a hormone with either systemic or reproductive functions; as growth factors for germ cell maintenance and in vitro culture; as a fertility control agent; remedy for sexual dysfunction or sex development disorders: diagnostics/treatment for ovarian tumors, such as serous adenocarcinoma, dysgerminoma, embryonal carcinoma,

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choriocarcinoma, teratoma, etc; The expression in thymus may indicate its utilities in T-cell development and thus its applications in immune related medical conditions, such as infection, allergy, immune deficiency, tissue/organ transplantation, etc.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 45

The translation product of this gene shares sequence homology with cytochrome C oxidase which is thought to be important in metabolic function of cells. This gene has now recently been published as estrogen response gene. See Genbank accession no. AB007618 and Mol. Cell. Biol. 18 (1), 442-449 (1998). See also J Immunol. Mar 1:154(5): 2384-2392 (1995), where the mouse homologue was published and implicated in siliocis.

This gene is expressed primarily in adipose tissue, kidney and fetal brain and to a lesser extent in several other tissues and organs.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolic diseases involving especially adipose tissue, brain and kidney. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the CNS and vascular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adipose tissue, kidney, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:169 as residues: Thr-5 to Ser-14.

The tissue distribution and homology to cytochrome C oxidase, estrogen response gene product and siliocis related gene product indicates that the protein products of this gene are useful for diagnosis and treatment of metabolic disorders in the CNS, adipose tissue and kidney, particularly siliocis.

FEATURES OF PROTEIN ENCODED BY GENE NO: 46

The translation product of this gene shares sequence homology with reticulocalbin. See, for example, J. Biochem. 117 (5), 1113-1119 (1995). Based on the

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sequence similarity, the translation product of this gene is expected to share certain biological activities with reticulocalbin, e.g., Ca++ binding activities. This gene product is sometimes hereinafter referred to as "Reticulocalbin-2".

This gene is expressed primarily in breast, endothelial cells, synovial, heart and smooth muscle cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the breast, vascular and skeletal/cardiac muscular system. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the breast, vascular and skeleto-muscular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., mammary tissue, endothelial cells, synovial tissue, heart and other cardiovascular tissue, and smooth muscle, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:170 as residues: Gly-16 to Arg-32, Ala-42 to Asn-50, Glu-66 to Gln-76, Arg-85 to Gly-94, Thr-108 to Asp-115, Trp-121 to Gly-130. Leu-137 to His-144, Glu-155 to Lys-161. Asp-175 to Ser-180, Glu-209 to Gly-217, Glu-232 to Glu-237, Thr-243 to Asp-261, Glu-287 to Arg-295.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis and treatment of diseases of the vascular and skeletal/cardiac muscular system. The homology of the gene with reticulocalbin indicates its biological function in regulating calcium store, a particularly important function in muscular cell types. The gene expression in the heart may indicate its utilities in diagnosis and remedy in heart failure, ischemic heart diseases, cardiomyopathy, hypertension, arrhythmia, etc. The abundant expression in the breast may indicate its applications in breast neoplasia and breast cancers, such as fibroadenoma, papillary carcinoma, ductal carcinoma, Pagetís disease, medullary carcinoma, mucinous carcinoma, tubular carcinoma, secretory carcinoma and apocrine carcinoma; juvenile hypertrophy and gynecomastia, mastitis and abscess, duct ectasia, fat necrosis and fibrocystic diseases, etc.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 47

The translation product of this gene shares weak sequence homology with H+-transporting ATP synthase which is thought to be important in cell metabolism or signal transduction.

This gene is expressed only in testis.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of some types of diseases and conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the brain and hematopoietic tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., testes and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

Since only one out of about a million expressed sequence tag is found in testes indicates that its expression is low and selectively in testes. Since some of the genes only expressed in testes are usually expressed in brain or in certain induced hematopoietic cells/tissues, it is speculated that this gene to be expressed in brain or hematopoietic cells/tissues and is useful for diagnosis and treatment of disorders these systems.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 48

The translation product of this gene shares sequence homology with human polymeric immunoglobulin receptor (accession No.X73079) which is thought to be important in antibody recognition and immune defenses. In one embodiment, polypeptides of the invention comprise the sequence GWYWCG (SEQ ID NO:285). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in placenta and to a lesser extent in corpus callosum and fetal liver and spleen.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, disorders of the immune system, e.g. autoimmune diseases and immunodeficiency. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, liver, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:172 as residues: Tyr-37 to Cys-49, Gly-51 to Tyr-56, Lys-88 to Trp-93, Leu-130 to Glu-136.

The tissue distribution and homology to human polymeric immunoglobulin receptor indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immune disorders, e.g. autoimmune diseases and immunodeficiencies.

FEATURES OF PROTEIN ENCODED BY GENE NO: 49

This gene is expressed in thymus.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune disorder. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., thymus and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immune disorders, e.g. autoimmunity and immunodeficiency.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 50

Preferred polypeptide encoded by this gene comprise the following amino acid sequence: MKVGARIRVKMSVNKAHPVVSTHWRWPAEWPQMFLHLAQEPRTE VKSRPLGLAGFIRQDSKTRKPLEQETIMSAADTALWPYGHGNREHQENELQKY LQYKDMHLLDSGQSLGHTHTLQGSHNLTALNI (SEQ ID NO:286). Polynucleotides encoding this polypeptide are also provided as are complementary polynucleotides thereto.

This gene is expressed primarily in adrenal gland, pituitary, T helper cells, and breast cells and to a lesser extent in a wide variety of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of the some diseases and conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adrenal gland, pituitary, T-cells and other blood cells, and mammary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:174 as residues: Gln-39 to Ser-47, Arg-57 to Glu-67, Tyr-82 to Gln-95.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of a wide range of disorders, such as immune and endocrine disorders.

30 FEATURES OF PROTEIN ENCODED BY GENE NO: 51

The translation product of this gene shares sequence homology with human Sop2p-like protein which is important in cytoskeleton structure. In one embodiment, polypeptides of the invention comprise the sequence SLHKNSVSQISVLSGGKAKCS QFCTTGMDGGMSIWDVKSLESALKDLKI (SEQ ID NO:287). Polynucleotides encoding this polypeptide are also encompassed by the invention. This gene maps to chromosome 7. Therefore, polynucleotides of the invention can be used in linkage

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analysis as a marker for chromosome 7.

This gene is expressed primarily in immune and hematopoietic tissues/cells and to a lesser extent in other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immunological and hematopoietic disorders and inflammation. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., immune and hematopoietic tissue/cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:175 as residues: Lys-49 to Gln-54, Ala-61 to Arg-66, Lys-82 to Lys-87, Glu-126 to Val-133, His-136 to Ile-141, Glu-175 to Ser-187, Asp-286 to Leu-296, Ala-298 to Ser-310.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immunological, hematopoietic, and inflammatory disorders, e.g, immunodeficiency, autoimmunity, inflammation.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 52

The translation product of this gene shares sequence homology with *Caenorhabditis elegans* R53.5 gene encoding a putative secreted protein without known function.

This gene is expressed primarily in endothelial cells, brain and several highly vascularized, and tumor tissues and to a lesser extent in other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, aberrant angiogensis and tumorigenesis. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes

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for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the vascular and brain system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, brain and other tissue of the nervous system, and vascular tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:176 as residues: Thr-43 to Asn-60, Thr-106 to Phe-115, Asp-122 to Arg-133, Arg-186 to Asp-192, Leu-211 to Lys-216.

The tissue distribution and homology to a *C. elegans* secreted protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis or treatment of disorders in vascular or brain system, e.g. aberrant angiogenesis, ischemia, neurodegeneration, etc.

FEATURES OF PROTEIN ENCODED BY GENE NO: 53

In one embodiment, polypeptides of the invention comprise the sequence EASKSSHAGLDLFSVAACHRF (SEQ ID NO:288). Polynucleotides encoding this polypeptide are also encompassed by the invention.

This gene is expressed primarily in T-cells and to a lesser extent in brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, lymphocytic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the lymphoid system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:177 as residues: Pro-3 to Thr-8, Arg-37 to Asp-46.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis, treatment, and cure of lymphocytic disorders.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 54

The translation product of this gene shares sequence homology with secreted cartilage matrix protein, a major component of the extracellular matrix of nonarticular cartilage which is thought to be important in cartilage structure. In specific embodiments, polypeptides of the invention comprise the sequence: RCKKCTEGPI DLVFVIDGSKSLGEENFEVVKQF (SEQ ID NO:297); VTGIIDSLTISPKAARVGL LQYSTQVH (SEQ ID NO:290); TEFTLRNFNSAKDMKKAVAHMKYM (SEQ ID NO:291): GKGSMTGLALKHMFERSFTQGEGARPF (SEQ ID NO:292); STRVP RAAIVFTDGRAQDDVSEWASKAKANGITMYAVGVGKAIE (SEQ ID NO:293); EELQEIASEPTNKHLFYAEDFSTMDEISEKLKKGICEALEDS (SEQ ID NO:294); TQRLEEMTQRM (SEQ ID NO:295); PQGCPEQPLH (SEQ ID NO:296); and/or YMGKGSMTGLALKHMFERSFT (SEQ ID NO:289). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in placenta, infant brain, prostate, fetal lung and to a lesser extent in endometrium and fetal tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormal placenta and pregnancy, disorder and injury in brain, prostate, and vasculature. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproduction, neuronal, and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, brain and other tissue of the nervous system, prostate, lung and endometrium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to cartilage matrix protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis,

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treatment, and cure of abnormalities in placenta and pregnancy, disorder and injury in brain, prostate, and vasculature.

FEATURES OF PROTEIN ENCODED BY GENE NO: 55

The translation product of this gene is the human ortholog of bovine and hamster CII-3, a succinate-ubiquinone oxidoreductase complex II membrane-intrinsic subunit, which is thought to be important in mitochondrial electron transport chain during metabolism. In specific embodiments, the polypeptides of the invention compriseMAALLLRHVGRHCLRAHFSPQLCIRNAVPLGTTAKEEMERFWNKNIG SNRPLSPHITIYS (SEQ ID NO:298); VFPLMYHTWNGIRHLMWDLGKGLKIPQL YQSG (SEQ ID NO:299); MAALLLRHVGRHCLRAH (SEQ ID NO:300); VKSLCL GPALIHTAKFAL (SEQ ID NO:301); VFPLMYHTWNGIRHLMWDLGKGL (SEQ ID NO:302).

This gene is expressed in 8-week old early stage human.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolism disorder. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the [insert system where a related disease state is likely, e.g., immune], expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis, treatment, and cure of metabolism disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 56

This gene is expressed primarily in umbilical vein endothelial cells, human ovarian tumor cells, human meningima cells, and human Jurkat membrane bound polysomes. In specific embodiments, polypeptides of the invention comprise the amino acid sequence: RVWDVRPFAPKERCVKIFQGNV (SEQ ID NO:303); HNFEKNLL

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RCSWSPDGSKIAAGSADRFVYV (SEQ ID NO:304); and/or WDTTSRRILYKLPG HAGSINEVAFHPDEPI (SEQ ID NO:305). Polynucleotides encoding these polypeptides are also encompassed by the invention.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation, immune and cardiovascular disorders and urogenital neoplasias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neurological, urogenital, reproductive system and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, cells, endothelial cells, ovary and other reproductive tissue, meningima, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:143 as residues: Phe-71 to Arg-76, Pro-82 to His-87, Glu-103 to Ala-111.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. In addition, expression in ovarian tumor cells suggests that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis, and treatment of ovarian tumors, and other tumors and neoplasias. Further, endothelial cell expression suggests a role in cadiovascular or respiratory/pulmonary disorders or infections (athsma, pulmonary edema, pneumonia).

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FEATURES OF PROTEIN ENCODED BY GENE NO: 57

The translation product of this gene shares sequence homology with type I collagen. In specific embodiments, the polypeptides of the invention comprise the sequence: GRIPAPAPSVPAGPDSR (SEQ ID NO:309); VRGRTVLRPGLDAEPE LSPE (SEQ ID NO:306); EQRVLERKLKKERKKEERQ (SEQ ID NO:307); ARRSG

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AELAWDYLCRWAQKHKNWRFQKTRQTWLLLHMYDSDKVPDEHFSTLLAYLE GLQGR (SEQ ID NO:255); and/or RLREAGLVAQHPP (SEQ ID NO:308).

Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in epididymus, prostate cell line (LNCAP), and pituitary gland; and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the epididymus, prostate (especially prostate cancer), and pituitary gland. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the male reproductive system and neuroendocrine system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., epididymus and other reproductive tissue, prostate, and pituitary gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to type I collagen, indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of the epididymus, prostate (especially prostate cancer), and pituitary gland.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 58

This gene is expressed primarily in the frontal cortex of the brain from a schizophrenic individual.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, schizophrenia. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of

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the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid-from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of schizophrenia.

FEATURES OF PROTEIN ENCODED BY GENE NO: 59

The polypeptide encoded by Gene 59 is homologous to human surface 4 integral membrane protein. In specific embodiments, the polypeptides of the invention comprise the sequence: TGCVLVLSRNFVQYACFGLFGIIALQTIAYSILWDLKF LMRN (SEQ ID NO:310); SRSEGKSMFAGVPTMRESSPKQYMQLGGRVLLV LMFMTLLHFDASFFSIVQNIVG (SEQ IDNO:311); GTAEDFADQFLRVTKQYLP HVARLCLISTFLEDGIRMFQWSEQRDYIDTTWNCGYLLAS (SEQ ID NO:312); LMRNESRS (SEQ ID NO:314); ASFLLSRTSWGTA (SEQ ID NO:315); and/or ASFLLSRTSWGTALMIL (SEQ ID NO:313). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in Hodgkin's lymphoma and lung; and to a lesser extent in many other human tissues.

Therefore, polynucleotides and polypeptides of the invention a suseful as reagents for differential identification of the tissue(s) or cell type is the seat in a biological sample and for diagnosis of diseases and conditions. This enclude, but are not limited to, Hodgkin's lymphoma, tumors or other abnormalities of the lung. Similarly, polypeptides and antibodies directed to these pulpoptides are weful in providing immunological probes for differential identification on of the page 3) or cell type(s). For a number of disorders of the above tissues of the Alis. z wid immune and respiratory systems, expression of this ge sign: No. 6 ther or C. Grin ue, and lower levels may be routinely detected in certain tissue pulmonary tissue, and cancerous and wounded tissue 200 :., serum, taken from plasma, urine, synovial fluid or spinal fluid) or anothe B 1 1 a level, i.e., an individual having such a disorder, relative to the id se aving the the expression level in healthy tissue or bodily fluid as income EQ ID disorder. Preferred epitopes include those comprising ::ie:::. NO:183 as residues: Met-20 to Trp-27.

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corresponding to this gene are useful for diagnosis and treatment of Hodgkin's lymphoma, tumors or other abnormalities of the lung.

FEATURES OF PROTEIN ENCODED BY GENE NO: 60

This gene is expressed primarily in bone cancer and stomach cancer, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, bone cancer and stomach cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the bone, and the stomach, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and stomach, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of bone cancer and stomach cancer and possibly other cancers.

FEATURES OF PROTEIN ENCODED BY GENE NO: 61

This gene is expressed primarily in epididymus, and lymph node of breast cancer, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the epididymus, and breast cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the epididymus and breast, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., epididymus and other reproductive tissue, lymphoid tissue, and mammary tissue, and cancerous and wounded tissues) or bodily fluids

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(e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:185 as residues: Arg-57 to Ser-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of the epididymus, and breast cancer.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 62

The translation product of this gene appears to be the human homolog of bovine NADH dehydrogenase which is thought to be important in cellular metabolism. In specific embodiments, the polypeptides of the invention comprise the amino acid sequence: SMSALTRLASFARVGGRLFRSGCARTAGDGGVRHAGGGVHIEPRY RQFPQLTRSQVFQSEFFSGLMWFWILWRFWHDSEEVLGHFPYPDPSQWTDEEL GIPPDDED (SEQ ID NO:323), or fragments thereof. Polynucleotides encoding this polypeptide are also encompassed by the invention.

This gene is expressed in larynx tumor, lymph node, brain amygdala, human cardiomyopathy, and retina.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases affecting cellular metabolism. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., larynx, lymphoid tissue, brain and other tissue of the nervous system, heart and cardiovascular tissue, and retina, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:208 as residues: Pro-27 to Gln-32, Arg-42 to Glu-51.

The tissue distribution and homology to NADH dehydrogenase indicates that

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polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of diseases involving cellular metabolism.

FEATURES OF PROTEIN ENCODED BY GENE NO: 63

This gene is expressed primarily in amygdala, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the amygdala. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the amygdala, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., amygdala, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:187 as residues: Gln-17 to Glu-29, Pro-41 to Phe-46, Ser-59 to Ile-70, Thr-97 to Leu-105.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of amygdala.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 64

This gene is expressed primarily in female bladder, and to a lesser extent in chronic synovitis and hemangiopericytoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, bladder cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the urinary tract, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bladder, synovial tissue, and

vascular tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:188 as residues: Pro-2 to Gln-7, Pro-27 to Phe-34.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatments of defects of the urinary tract, especially bladder cancer.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 65

This gene is expressed primarily in fetal spleen, and to a lesser extent in hemangiopericytoma, thymus, and synovial sarcoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, defects of immune of hematopoietic systems. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune of hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., spleen, vascular tissue, thymus, blood cells, and synovial tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The protein product of this gene is useful for treatment of defects of the immune or hematopoietic systems, because of the gene's expression in thymus and spleen.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 66

This gene is expressed primarily in human pituitary and to a lesser extent in placenta and fetal lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, endocrine growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pituitary and other endocrine tissue, placenta, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:190 as residues: Val-38 to Asn-44, Gly-53 to Ser-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment of growth disorders related to pituitary dysfunction.

FEATURES OF PROTEIN ENCODED BY GENE NO: 67

The translation product of this gene shares sequence homology with a *Caenorhabditis elegans* gene of unknown function. In specific embodiments, the polypeptides of the invention comprise the sequence: DPRRPNKVLRYKPPPSE CNPALDDPTP (SEQ ID NO:317): DYMNLLGMIFSMCGLMLKLKWCAWVA VYCS (SEQ ID NO:318); FISFANSRSSEDTKQMMSSF (SEQ ID NO:316); and/or MLSISAVVMSYLQNPQPMTPPW (SEQ ID NO:319). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in primary breast cancer and lymph node breast cancer and to a lesser extent in adult brain, lung cancer, colon cancer, epithelioid sarcoma, and Caco-2 cell line.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cancer and tumor tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, lymphoid tissue, brain and other tissue of the nervous system, lung, colon, and

epithelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:191 as residues: Asn-34 to Lys-42.

The tissue distribution in a variety of cancer tissues indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of a variety of cancer and tumor types.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 68

The translation product of this gene shares sequence homology with steroid membrane binding protein. The translation product of this gene has recently been published as progesterone binding protein. See Genbank AJ002030. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: AAGDGDVKLGTLGSGSESSNDGGSESPGDAGAAAXGGGWAAAALALLTG GGE (SEQ ID NO:320).

This gene is expressed primarily in breast, and to a lesser extent in placenta and fetal tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, breast cancer or developmental disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of breast or fetal tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, placenta, and fetal tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:192 as residues: Pro-43 to Asp-49, Gln-54 to Pro-64, Asp-110 to Asp-118, Lys-138 to Tyr-143, Pro-150 to Asp-170.

The tissue distribution and homology to steroid membrane binding protein and to progesterone binding protein indicates that the protein products of this gene are

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useful for treatment of breast cancers, especially those caused by estrogen and progesterone binding.

FEATURES OF PROTEIN ENCODED BY GENE NO: 69

Preferred polypeptides encoded by this gene comprise the following amino acid sequence: AADNYGIPRACRNSARSYGAAWLLLXPAGSSRVEPTQDISISDQLGG QDVPVFRNLSLLVVGVGAVFSLLFHLGTRERRRPHAXEPGEHTPLLAPATAQPL LLWKHWLREXAFYQVGILYMTTRLIVNLSQTYMAMYLTYSLHLPKKFIATIPLV MYLSGFLSSFLMKPINKCIGRN (SEQ ID NO:321).

This gene is expressed primarily in macrophage (GM-CSF treated), and to a lesser extent in monocytes and dendritic cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and infection. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., macrophages and other blood cells, and dendritic cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for treatment of infection or inflammation or other events or defects involving the immune system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 70

This gene is expressed primarily in adult brain and to a lesser extent in thyroid, 12 week old early stage human, and stromal cell TF274.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurological or neuro-endocrine diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes

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for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous or endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., brain and other tissue of the nervous system, thyroid, and stromal cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:194 as residues: Pro-65 to Cys-71.

The tissue distribution indicates that the protein products of this gene are useful for treatment and diagnosis of neurological diseases or metabolic conditions involving the neuro-endocrine system.

15 FEATURES OF PROTEIN ENCODED BY GENE NO: 71

This gene is expressed in T-cell helper and to a lesser extent in adult brain and adult testes.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune disorders, meningitis or reproductive problems. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neural and reproductive systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, brain and other tissue of the nervous system, testes and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:195 as residues: Val-18 to Tyr-24, Ala-89 to Asp-99, Asp-104 to Ala-117, Leu-121 to Pro-136.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis immune and

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reproductive disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 72

The translated polypeptide of this contig has a high degree of identity with the Ob Receptor-Associated Protein deposited as GenBank Accession No. 2266638. No function has been determined for the Ob Receptor-Associated Protein, however it is expressed upon stimulation of the Ob Receptor by Leptin.

This gene is expressed in T-cells and to a lesser extent in endothelial and bone marrow cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, acute lymphoblastic leukemia, hematapoetic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematapoetic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, endothelial cells, and bone marrow, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO: 196 as residues: Ser-61 to Trp-70.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of leukemia and other disorders of the primary immune system. In addition, since this gene appears to be related to the Ob Receptor-Related Protein, it is likely that this polypeptide is also involved in the Ob/Leptin signal transduction cascade. As a result, this protein may be of use in the molecular diagnosis and therapeutic intervention of obesity and related disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 73

The translation product of this contig has homology with furin, a protein thought to be a key endopeptidase in the constitutive secretory pathway. The identification and initial characterization of Furin was reported by Takahasi and

colleagues (Biochem Biophys Res Commun 1993 Sep 15:195(2):1019-1026).

This gene is expressed in neutrophils.

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Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the immune system such as allergies, wound healing and antigen recognition. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., neutrophils and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment of allergies or other immune disorders since neutrophils are an important part of an allergic response. Further, since this protein appears to be related to Furin, it can be used diagnostically and therapeutically to treat secretory protein processing disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 74

This gene is expressed in the frontal cortex.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, of the motor activity and sensory functions that involve the central nervous system. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene

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expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the detection and treatment of neural disorders that affect cognitive functions.

FEATURES OF PROTEIN ENCODED BY GENE NO: 75

The translation product of this gene shares sequence homology with inorganic pyrophophatase which is thought to be important in the catalysis the hydrolysis of diphosphate bonds, chiefly in nucleoside di- and triphosphates and essential enzymes that are important for controlling the cellular levels of inorganic pyrophosphate (PPi). The bovine homolog of this gene has been identified by Yang and Wensel (J. Biol. Chem. 267:24641-24647 (1992)).

This gene is expressed in osteoclastoma cells and to a lesser extent in epithelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoporosis and other bone weakening diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the skeletal system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., bone, and epithelial cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:199 as residues: Lys-22 to Tyr-28, Asp-64 to Lys-77, Pro-86 to Ile-91, Gln-99 to Pro-119, Tyr-169 to Asp-174, Lys-176 to Gly-181, Trp-189 to Asn-202, Lys-233 to Gly-239, Ser-250 to Asp-257.

The tissue distribution and homology to inorganic pyrophophatase indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of osteoporosis through the removal of bone by demineralization.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 76

The translation product of this gene shares exact sequence homology with ATP sulfurylase/APS kinase (GenBank Accession No. 2673862) which is thought to be important in biosynthesis of the activated sulfate donor, adenosine 3'-phosphate 5'-phosphosulfate, involves the sequential action of two enzyme activities: ATP sulfurylase, which catalyzes the formation of adenosine 5'-phosphosulfate (APS) from ATP and free sulfate, and APS kinase, which subsequently phosphorylates APS to produce adenosine 3'-phosphate 5'-phosphosulfate.

This gene is expressed in osteoclastoma cells and to a lesser extent in developmental tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, antibiotic resistant bacterial infections, osteoarthritis and other auto immune diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune or skeletal structure expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and developmental tissues, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:200 as residues: Asn-15 to Trp-20, Ser-36 to Gly-41, Pro-103 to Val-110, Pro-134 to Arg-143, Leu-173 to Arg-178, Ser-190 to Ala-197, His-314 to Arg-319, Arg-354 to Asn-362, Asp-391 to Arg-397, Glu-402 to Asp-409. Asp-434 to Leu-439, Glu-441 to Arg-446, Gly-455 to Asp-462, Pro-528 to His-541, Asn-566 to Arg-571, Tyr-574 to Glu-581, Thr-589 to Glu-603.

The tissue distribution and homology to ATP sulfurylase/APS kinase indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment or detection of autoimmune diseases.

FEATURES OF PROTEIN ENCODED BY GENE NO: 77

This polypeptide is identical to the SLP-76-associated protein reported by Musci and colleagues (J. Biol. Chem. 272 (18), 11674-11677 (1997)) and to the FYB protein

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reported by da Silva and coworkers (Proc. Natl. Acad. Sci. U.S.A. (1997) In press). These proteins have been reported to be novel T-cell Proteins which bind FYN and SLP-76 and regulate IL-2 production. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: RITDNPEGKWLGRTARGSYGYIK TTAVEIXYDSLKLKKDSLGAPSRPIEDDQEVYDDVAEQDDISSHSQSGGGIFPP PPDDDIYDGIEEEDADDGFPAPPKQLDMGDEVYDDVDTSDFPVSSAEMSQGTNV GKAKTEEKDLKKLKKQXKEXKDFRKKFKYDGEIRVLYSTKVTTSITSKKWGT RDLQVKPGESLEVIQTTDDTKVLCRNEEGKYGYVLRSYLADNDGEIYDDIADGC IYDND (SEQ ID NO:322).

This gene is expressed in CD34 positive cells (hematopoietic progenitor cells) and to a lesser extent in adult spleen derived from a chronic lymphocytic leukemia patient.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are 15 not limited to, chronic lymphocytic leukemia; hematopoietic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels 20 may be routinely detected in certain tissues (e.g., T-cells and other blood cells, bone marrow, hematopoietic cells, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily 25 fluid from an individual not having the disorder. Further, nucleic acids and polypeptides of the present invention are useful both diagnostically and therapeutically in the intervention of immune and other disorders in which the ability to alter IL-2 expression is desired. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:201 as residues: Ala-17 to Lys-37, Val-39 to Ser-45, Lys-59 to His-70, 30 Arg-90 to Leu-95, Lys-97 to Lys-107, Ser-117 to Leu-124, Phe-133 to Ser-138, Trp-146 to Leu-167, Pro-175 to Asn-185, Lys-190 to Ser-211, Pro-213 to Ser-222, His-230 to Pro-235, Pro-240 to Pro-246, Pro-253 to Gly-261, Leu-271 to Leu-303, Leu-305 to Leu-326, Lys-343 to Leu-349, Thr-363 to Leu-371, Arg-373 to Tyr-381, Tyr-391 to Leu-401, Pro-404 to Val-414, Ser-426 to Ser-432, Ile-448 to Ser-457, Gln-462 35 to Trp-468, Lys-477 to Ser-501, Asp-518 to Ser-523, Ala-541 to Gln-554.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment of a variety of hematopoietic disorders. The noted expression of this gene in the hematopoietic progenitor cell compartment - as determined by its expression on CD34 positive hematopoietic stem and progenitor cells - indicates that it plays a critical role in the expansion or proliferation of hematopoietic stem/progenitor cells, as well as in the differentiation of the various blood cell lineages. Thus it could be useful in the reconstitution of the hematopoietic system of patients with leukemias and other hematopoietic diseases.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 78

This gene is homologous to heparin cofactor II (HCII) which is a 66-kDa plasma glycoprotein that inhibits thrombin rapidly in the presence of dermatan sulfate or heparin.

This gene is expressed in apoptotic and anergic T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, thrombopienia T-cell lymphomas: Hodgkin's lymphoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system most notably the T-cell compartment, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The homology to heparin cofactor II (HCII) and the tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of hematopoietic disorders particularly in thrombopoesis, most notably of the T-cell compartment. This could include immune modulation, inflammation, immune surveillance, graft rejection, and autoimmunity.

35 FEATURES OF PROTEIN ENCODED BY GENE NO: 79

The translation product of this gene shares sequence homology with a mouse

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protein believed to represent an integral membrane protein.

This gene is expressed in fetal cochlea and epididymus and to a lesser extent in adult spleen and osteoclastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoclastoma; disorders of the inner ear; male fertility disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the inner ear; male reproductive tract; bone; and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., cochlea, epididymus and other reproductive tissue, spleen, and bone, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:203 as residues: Lys-13 to Gly-23, Cys-38 to Asp-43, Gly-48 to Trp-53, Cys-223 to Ile-237, Ile-240 to Ser-246.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment of hearing and fertility disorders. Likewise, it may have a role in the modulation of immune function and in the treatment of osteoporosis.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 80

The translation product of this gene shares sequence homology with reticulocalbin which is thought to be important in the binding of calcium, particularly within the endoplasmic reticulum.

This gene is expressed in endothelial cells and stromal cells and to a lesser extent in osteoblasts, osteoclasts, and T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoperosis; osteoclastomas; T-cell lymphomas; Hodgkin's disease. Similarly, polypeptides and antibodies directed to these polypeptides are useful in

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providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the vasculature, bone, and immune systems - particularly the T-cell compartments, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, stromal cells, bone, T-cells and other blood cells, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:204 as residues: Lys-20 to Arg-27, Pro-32 to Asp-48, Leu-64 to Arg-72, Asp-108 to Lys-114, Glu-128 to Thr-133, Asp-139 to Phe-147. Thr-196 to Ala-204, Tyr-218 to Glu-228, Val-230 to Gln-236, Arg-241 to Lys-255, Glu-276 to Lys-287.

The tissue distribution and homology to reticulocalbin indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of bone disorders such as osteoporosis; the diagnosis and treatment of T-cell lymphomas and Hodgkin's lymphoma; and the treatment of diseases and defects of the vasculature, such as vascular leak syndrome and aberrant angiogenesis that accompanies tumor growth.

FEATURES OF PROTEIN ENCODED BY GENE NO: 81

The translation product of this gene shares sequence homology with a family of peptide transport genes - particularly the AtPTR2-B gene from *Arabidopsis* - which are thought to be important in the uptake of small peptides.

This gene is expressed in a number of fetal tissues, most notably lung, brain, cochlea, and liver/spleen, and to a lesser extent in osteoclastoma and endometrial tumors.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoclastoma; endometrial tumors; cancer; leukemias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the bone and endometrium, expression of this gene at significantly higher or lower levels may be

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routinely detected in certain tissues (e.g., fetal tissue, pulmonary tissue, bone, brain and other tissue of the nervous system, cochlea, liver, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:205 as residues: Lys-186 to Asn-199, Pro-202 to Ala-207.

The tissue distribution and homology to peptide transport genes indicates that polynucleotides and polypeptides corresponding to this gene are useful for the control of cell proliferation, owing to its strong expression in fetal tissues undergoing active cell division, as well as its expression in a variety of tumors or cancers of adult tissues. Potentially, it may regulate the uptake of peptides that stimulate cell proliferation. This gene product may also be useful in stimulating the uptake of a variety of peptide-based drug compounds.

FEATURES OF PROTEIN ENCODED BY GENE NO: 82

This gene is expressed in fetal liver and spleen and to a lesser extent in endothelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer and tumors of a hematopoietic and/or endothelial cell origin; leukemias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system and/or vasculature, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., liver, spleen, endothelial cells, vascular tissue, and tissue and cells of the immune system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:206 as residues: Met-1 to Asp-9, Arg-66 to Gly-76, Asp-164 to Arg-171.

The tissue distribution indicates that polynucleotides and polypeptides

corresponding to this gene are useful for the treatment of disorders of the immune system. Expression of this gene product in both fetal liver/spleen and endothelial cells indicates that it may be expressed in the hemangioblast, the progenitor cell for both the immune system and the vasculature. Thus, it is most likely expressed in hematopoietic stem cells, and may be useful for the expansion of hematopoietic stem and progenitor cells in conjunction with cancer treatment for a variety of leukemias.

FEATURES OF PROTEIN ENCODED BY GENE NO: 84

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The translation product of this gene shares sequence homology with NADH dehydrogenase which is thought to be important in cellular metabolism.

This gene is expressed in fetal dura mater and to a lesser extent in T-cells and hypothalamus.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases affecting cellular metabolism. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., fetal tissue. T-cells and other blood cells, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:208 as residues: Pro-27 to Gln-32, Arg-42 to Glu-51.

The tissue distribution and homology to NADH dehydrogenase indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of diseases involving cellular metabolism.

FEATURES OF PROTEIN ENCODED BY GENE NO: 85

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The translation product of this gene shares sequence homology with I-TRAF, a novel TNF receptor associated factor (TRAF)-interacting protein that regulates TNF receptor-mediated signal transduction. This protein is thought to be important in regulating the cellular response to tumor necrosis factor (TNF), which is an important mediator of inflammation.

This gene is expressed in endothelial cells and to a lesser extent in glioblastoma and osteoblastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation; glioblastoma and osteoblastoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, bone, and glial cells and tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:209 as residues: Glu-15 to Thr-22, Glu-46 to Leu-62, Arg-103 to Glu-119, Gln-127 to Glu-132, Asn-152 to Trp-158, Gln-191 to Gln-210, Glu-264 to Thr-271, Tyr-282 to Leu-288, Trp-319 to Thr-331, Glu-335 to Ser-348, Ser-353 to Ser-358, Asp-382 to Asn-392.

The tissue distribution and homology to I-TRAF indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of inflammatory diseases, including rheumatoid arthritis, sepsis, inflammatory bowel disease, and psoriasis, particularly where tumor necrosis factor is known to be involved.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 86

This gene has homology with a candidate gene involved in X-linked Retinopathy reported by Wong and colleagues (Genomics 15:467-471 (1993)).

This gene is expressed in a T-cell line.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and autoimmune diseases; T-cell lymphoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of inflammatory disorders such as sepsis, inflammatory bowel disease, psoriasis, and rheumatoid arthritis as well as autoimmune disease such as lupus. It could also be useful in immune modulation and in the process of immune surveillance. The present invention can be used diagnostically and therapeutically to treat X-linked Retinopathy.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 87

This gene is expressed in human brain tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, brain disorders; neurodegenerative disorders; tumors of a brain origin. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma,

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urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:211 as residues: Cys-32 to Tyr-38.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of CNS disorders such as epilepsy, paranoia, depression, Alzheimer's disease, and schizophrenia. It could be useful in the survival and/or proliferation of neurons and could effect neuronal regeneration.

Last AA of ORF	30	44	69	∞	38	22	109
First AA of Secreted Portion		27	45	26	. 25	23	21
Last AA of Sig Pep		26	44	25	24	22	20
First AA of Sig Pep	1	1		-	_		_
SEQ SEQ ×	125	126	212	213	127	128	129
5' NT AA First Last of AA of D of Signal NO: Sign Sign Pep A Pep Pep	353	128	170	413	66	006	103
of of Start Codo	353	128	170	413	66	006	103
3' NT of Clone Seq.	1607	1786	1487	1637	1212	2061	733
S' NT 3' NT of Of Clone Clone Seq.	247	87	79	394		882	01
Total NT Seq.	1679	1830	1487	1653	1212	2061	1412
X SEQ	Ξ	12	86	66	13	4	15
Vector	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071	xxxxx 03/19/98	209641	97923 03/07/97 209071	97923 03/07/97 209071	97923 97923 03/07/97 209071 05/22/97
cDNA Clone ID	HAGEW82	HAGFY16	HBMCF37	HFLQB16	HALAA60	HAPBL78	HASAV70
Gene No.	_	2	2	2	3	4	2

Last AA of ORF	62	29	52	56	215	48
Last First AA of of Sig Secreted Pep Portion (18		24		19	. 27
Last AA of Sig Pep	17		23	17	18	26
AAA of Sig		-		_	-	-
AA SEQ ID NO: Y	130	131	132	133	134	135
S' NT of AA F First SEQ AA of ID Signal NO: Pep Y I	538	181	. 98	192	401	793
of of Start	538	181	98	192	401	793
3' NT of Clone Seq.	088	683	1007	1393	1070	2011
S' NT 3' NT of of Clone Clone Seq.	276		98	132	277	614
Total NT Seq.	1052	683	1054	1393	1215	2042
× Š B Š K	16	17	18		20	21
Vector	Uni-ZAP XR					
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/93	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/93	97923 03/07/97 209071 05/22/97
cDNA Clone ID	HBNAF22	HBNBL77	HCDDR90	HCEEF50	HCEMU42	HCENE16
Gene No.	9	7	∞ -	6	10	=

, D	67	51	539	08	56	48	200
rA ed	24	30	31	23	27	37	28
Last AA of Sig Pep	23	59	30	22	26	36	27
First AA of Sig Pep	_	-		_	_		_
AA SEQ ID NO:	136	137	138	214	139	215	140
5' NT of AA First SEQ AA of ID Signal NO: Pep Y	69	68	808	515	961	295	70
5° NT of Start Codon	69	68	808	515	961	295	70
	1872	289	3532	1115	907	734	717
s' NT 3' NT of of Clone Clone Seq. Seq.	21	_	2821	435	171	25	_
Total NT Seq.	1872	289	3533	1145	1148	734	717
× S B S X	22	23	24	001	25	101	26
Vector	Uni-ZAP XR	ZAP Express	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/93	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/27/97	97923 03/07/97 209071	209179 07/24/97	97923 03/07/97 209071 05/27/97	97923 03/07/97 209071 05/22/97
cDNA Clone ID	HMSJJ74	HCUBF15	HE2DE47	HE2DE47	HKMLH01	HE6DG34	HE9DG49
Gene No.	13	13	14	4.	15	15	91

Last AA of ORF	202	215	185	101		61
	2	7	_			
First AA of Secreted Portion	29	23	26	43	31	
Last AA of Sig Pep	28	22	25	42	30	
First AA of Sig Pep		1	1			_
AA SEQ ID NÖ:	216	141	217	142	143	144
S'NT AA First Last AA of ID of of Signal NO: Sign Sign Sign Sign Sign Sign Sign Sign	78	38	149	128	294	496
of of Star	78	38	149	128	294	496
3' NT of Clone Seq.	713	1099	1080	941	756	2093
S' NT 3' NT of of Clone Clone Seq. Seq.	17	_	_	171	62	408
Total NT Seq.	713	6601	1080	941	756	2100
SEQ SEQ X Sign	102	27	103	28	29	30
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 (15/27/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/93
cDNA Clone ID	HE9DG49	HELBA06	HELBA06	HSLFM29	HELBW38	HETHN28
Gene No.	16	17	17	18	61	20

Last AA of ORF	29	», , , , , , , , , , , , , , , , , , ,	7	38	130	31	13
First AA Lof Of A Secreted Portion C		67		. 17	27	22	
Last AA I of Sig		78		91	26	21	
First AA of Sig Pep	_	_	_	-	1	_	
AA SEQ ID YOO:	145	146	147	148	149.	150	151
5' NT of First AA of Signal I	567	21	210	242	178	144	1104
5' NT of Start Codon	567	21	210	242	178	144	1104
3' NT of Clone Seq.	1392	409	1322	710	1161	938	1581
S' NT 3' NT of of Clone Clone Seq. Seq.	475		_	-	110		974
Total NT Seq.	1448	456	1326	710	1188	956	1603
X S B S X	31	32	33	34	35	36	37
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/93	03/07/20 03/07/97 209071	97923 03/07/97 209071	97923 03/07/97 209071 05/27/97	97923 03/07/97 209071 05/22/97	97924 03/07/97
cDNA Clone ID	HFCDK17	HFEAF41	HFKFL13	HFSBG13	HFTBE43	HFTDJ36	HKTAC77
Gene No.	21	22	23	24	25	26	27

Last AA of ORF	_	19	52	194	8	30	68	68	% 2	C [13/	47	4
First AA Loft		33		33		31	20	. 23	61	17	21	28	78
Last AA F of Sig S		32		32	18	30	61	22	18	07	70	27	27
First AA of Sig Pep	_		_		I		_	1	1	-	_	-	_
SEQ ₹ÖÖÖ.≻	152	153	154	155	156	157	158	218	159	160	219	220	161
5' NT of First AA of Signal	209	119	581	126	43	171	55	58	17	15	72	54	269
5' NT of Start Codon		119	581	126	43	171	55	58	17	15	72	54	269
	1067	629	1793	1123	875	843	489	489	534	1374	640	1399	596
S' NT 3' NT of Clone Clone Seq. Seq.	55	-	408	13	-	_	3	9		_	28	40	-
Total C NT Seq.	6801	629	1964	1522	875	843	489	489	534	1374	640	1529	296
×ÖBÖX	38	39	40	41	42	43	44	104	45	46	105	901	47
Vector	pBluescript	pBluescript	Lambda ZAP II	Uni-ZAP XR	Uni-ZAP XR	pSport1	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97924	97924 03/07/97	97924	97924	97924	97924	97924	97924	97924 03/07/97	97924 03/07/97	97924	97924	97924 03/07/97
cDNA Clone ID		HLHSV96	HLQBQ86	HLTBX31	HLTCJ63	HMKAH44	HMQAJ64	НМQАЈ64	HOABG65	HODCL36	HODCL36	HODCL36	HODCL50
Gene No.	28	29	30	31	32	33	34	34	35	36	36	36	37

Last AA of	JY CK	77	60	322	69	319	82	30	71	280	7.5	22	326	183
First AA of Secreted	Portion	G	8 8	20	32	19	22		61 .	31	31		20	24
Last AA of Sig	rep	ŗ	/-	19	31	09	21		18	30	30		19	23
First AA of Sig	rep	-	-	-		-	-			-	-	_		-
AA SEQ UÖ:	Y	701	163	<u>2</u>	221	165	222	991	167	891	223	691	170	224
S' NT OF AA OF EICST SEQ AA OF ID Signal NO:	Pep	1/0	638	66	928	150	239	432	142	25	433	217	57	35
	_	1/0	638	99	928	150	239	432	142	25	433	217	57	35
3' NT of Clone Seq.	CCO	877	2020	2432	2435	2340	791	601	337	1141	1166	1148	809	586
S' NT3' NT of of Clone Clone Seq. Seq.	5			848	849	1627	92	188	-	_	21	63	164	4
	Seq.	108	2020	2432	2435	2340	805	601	359	1141	9911	0951	1507	286
SEQUE:	×	48	49	50	107	51	108	52	53	54	601	55	56	110
	Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pBluescript SK-	pBluescript SK-	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and	Date	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	979 <u>2</u> 4 03/07/97	97924 03/07/97	97924	97924 03/07/97	97924 03/07/97
cDNA	Clone ID	HODCV74	HODCZ16	HTOEU03	HTOEU03	HPBCJ74	HPBCJ74	HPMBU33	HSAUL66	HSIDQ18	HSIDQ18	HSJBB37	HSJBQ79	HSJBQ79
Gene	No.	38	39	40	40	41	4	45	43	44	44	45	46	46

Last AA of ORF	89	158	70	122	128	6	371
First AA of Secreted Portion	36	16	20	61	31		2
	35	15	61	18	30		-
First Last AA AA of of Sig Sig Pep Pep	-		-			_	
AA SEQ ID NO: Y	171	172	225	173	174	226	521
5' NT of First AA of Signal Pep	83	163	155	115	52	829	114
S' NT of Start Codon	83	163	155	115	52	829	114
3' NT' of Clone Seq.	450	1147	1134	777	598	1333	1554
S' NT 3' NT of of Clone Clone Seq. Seq.	_	-	_		48	594	443
Total NT Seq.	450	1147	1134	777	1611	1333	1580
NT SEQ BD NÖ:	57	58	Ξ	59	09	112	19
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pBluescript.	pBluescript	Uni-ZAP XR
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	209235
cDNA Clone ID	HTEGA76	HTEJN13	HTEJN13	HTHBL86	HTSF071	HTSF071	HAPNO80
Gene No.	47	48	48	49	50	50	51

Last AA of ORF	137	215	54	22	102	47
First AA of Secreted Portion	29	56	33	21	34	39
Last AA of Sig Pep	28	28	32	20	33	38
First AA of Sig Pep	_	-		_		
AA SEQ ID NO: Y	227	176	177	178	179	180
Signal NO: Sig Sig NY Pep Pep	244	182	16	150	231	703
S' NT of Start	244	182	97	150	231	703
3' NT of Clone Seq.	708	1034	361	1638	1303	1011
5' NT 3' NT of of Clone Clone Seq. Seq.	249	105	_	-	35	655
Total NT Seq.	1015	1117	361	1668	1353	1011
× SeQ	113	62	63	64	92	99
Vector	Uni-ZAP XR	pBluescript	pSport!	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97
cDNA Clone ID	HAUCC47	HBMCL41	HCFLD84	НЕ8ЕМ69	HE8EZ48	HEBGF73
Gene No.	51	52	53	54	55	56

Last AA of ORF	95	94	26	01	64	21
Last First AA I of of Sig Secreted Pep Portion (36	30	22		20	22
Last AA of Sig Pep	35	29	21		19	21
AAA of Sig Pep	1	-				
SEQ NO:	181	182	183	184	185	186
NT of rrst gnal gnal ep	459	63	839	270	272	127
S' NT of Start Codon	459	63	839	270	272	127
3' NT of Clone Seq.	0601	960	1851	711	935	484
S' NT 3' NT of of Clone Clone Seq.	267	_	765	∞	=	113
Total NT Seq.	1193	960	1657	711	935	504
SEQ NÖ:	19	89	69	70	71	72
Vector	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Lambda ZAP II	Lambda ZAP II	Lambda ZAP II
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97
cDNA Clone ID	HFEBF41	HFRBU14	HFVGZ79	HHGCM76	ННССО88	HHGCP52
Gene No.	57	58	59	09	19	62

Last AA of ORF	131	89	44	49	22	169
First AA of Secreted Portion	19	33	28	37	12	15
Last AA of Sig Pep	18	32	27	36		41
irst AA of Sig Pep	-				_	-
AA SEQ ID NÖ:	187	188	681	061	228	192
5' NT of AA I First SEQ AA of ID Signal NO: Pep Y	96	248	630	167	575	187
s' NT of Start Codon	96	248	630	191		187
3' NT of Clone Seq.	620	581	1786	800	1076	1888
S' NT 3' NT of of Clone Clone Seq.	_	156	537	116	398	18
Total NT Seq.	620	581	1843	1441	1076	2776
× SEQ	73	74	75	76	114	78
Vector	Lambda ZAP II	Lambda ZAP II	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pCMVSport 3.0
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/27/97	97958 03/13/97 209072	97958 03/13/97 209072	97958 03/13/97 209072 05/2/97	97958 03/13/97 209072 05/22/97
cDNA Clone ID	HHGDB72	HHGDI71	HHSDI45	HHSEB66	HJPAZ83	HLDBO49
Gene No.	63	64	65	99	<i>L</i> 9	89

Last AA of ORF	65	131	16	175	69	24	72
First AA of Secreted Portion	23	23	33	24	. 27	21	26
	22	22	32	23	26	20	25
First Last AA AA of of of Sig Sig Pep	-	_]	_	•		ed	
AA SEQ ID NO: Y	193	229	194	561	196	197	198
S' NT AA F F Signal NO: Signal NO: Pep Y H	534	534	40	238	286	28	14
of of Start	534	534	40	238	286	28	4
3' NT of Clone Seq.	1480	1487	1077	780	770	481	623
South Seq. Seq. Seq. Seq. Seq.	401	401	33	81	101	_	
Total NT Seq.	1525	1487	1563	1020	770	481	644
× SEQ	79	115	80	18	82	83	84
Vector	pCMVSport 3.0	pCMVSport 3.0	Uni-ZAP XR	Uni-Zap XR	Uni-ZAP XR	Uni-ZAP XR	pCMVSport 3.0
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	209226 08/28/97	97958 03/13/97 209072 05/22/97	97957 03/13/97 209073 05/27/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HLDBQ19	НГДВО19	HMSGT42	HMWIC78	HTTCT79	HNGJU84	HNTAC73
Gene No.	69	69	70	71	72	73	74

Last AA of ORF	288	27	623	09	648	28
rA on	13		31	33	31	22
	12		30	32	30	21
rst f f gp		_	-	-		_
AA SEQ NÖ:	661	230	200	231	201	232
S NT 3' NT of AA Fin of AA Fin of AA Fin of AA Glone Clone of AA of ID of Start Seq. Seq. Seq. Start Signal NO: Signal Seq. Y Pe	86	545	99	477	251	677
5' NT of Start Codon	86		2 6	477	251	677
3' NT of Clone Seq.	1284	1283	1747	1747	2566	1098
S' NT of Clone Seq.	435	428	290	288	1843 2566	375
Total NT Seq.	1351	1350	2527	2527	2566	8601
× Š B Š Š	85	911	98	117	87	811
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAF XR
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/27/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HOSEI45	JIOSE145	HOSFD58	HOSFD58	HSAUM95	HSAUM95
Gene No.	75	75	76	76	77	77

Last AA of ORF	54	265	17	314	206	194
Last of Of AA Sig Secreted Of Pep Portion ORF	33	12			21	70
Last AA I of Sig Pep	32	11		19	20	69
irst AA of Sig Pep	1	I	_	_	_	-
AA SEQ NÖ:	202	203	233	204	205	206
S' NT of AA F First SEQ AA of ID Signal NO:	83	188	315	92	414	157
S NT3' NT of of S' NT F of Of S' NT F F Otal Clone Clone of A NT Seq. Seq. Start Seq.	83	188	315	92	414	157
3' NT of Clone Seq.	540	1165	1166	2449	2058	1411
5' NT: of Clone Seq.	_	152	152	1149	476	345
Total NT Seq.	540	1863	1679	2478	2058	1411
SEQ X S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.	88	68	611	06	91	92
Vector	Uni-ZAP XR	pBluescript				
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/27/97	97957 03/13/97 209073 05/27/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HSAUR67	HSKD181	HSKDI81	HSKDW91	HTLEX50	HSKHL65
Gene No.	78	79	79	80	81	82

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•	87
	TS 16 12
AA AA 71 71 71 329	6 1 2
s s s s s s s s s s s s s s s s s s s	2 2 3
First AA Secreted Portion 38	- R - R
AA Find Signature 197 37 37 37 37 37 37 37 37 37 37 37 37 37	6 2
First Last AA	7 500 500 500 500 500 500 500 500 500 50
ZOT	33
A S Z C	223 445 223
S'NT of First AA of Signal Pep 526	
S of Start Start S Codon S 11 526 11 526 184 397	22 4 4 22
NT 5. one 6. eq. Co. 2184	2063
30 %	48 224
of Seq. Seq. 345	394
S C S S C S C S C S C S C S C S C S C S	27 27 878
248	12 8 12 83 1
ZSZ	
Vector	Uni-ZAP XR Uni-ZAP XR Uni-ZAP XF 17 2097 17 2097 1877 1877 Uni-ZAP 1877 Uni-ZAP 1877 Uni-ZAP 1877 Uni-ZAP 1877 Uni-ZAP
Vector Bluescri	
37.0	05/22/97 97957 03/13/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97 05/22/97
ATCC Deposit Nr and Date 97957 03/13/9	22/20/20/20/20/20/20/20/20/20/20/20/20/2
AUZ 6.00	HFGA11 97/200 2005/2005/2005/2005/2005/2005/2005/
cDNA Clone ID HSKHL65	HHFGAII HHFGAII HWTBLA 86 HCAC
cDNA Clone IC	臣严于一
	88 83 83 83 83 83 83 83 83 83 83 83 83 8
Ogene No.	

						-	-		C 17					*
Gene No.	cDNA Clone ID	ATCC Deposit Nr and Date	Vector	SEQ X	Total NT Seq.	S' NT 3' NT of of of S' NT Clone Clone of NT Seq. Seq. Start Seq.	of Clone Seq.	5' NT of Start Codon	of of First AA of Signal Pep	SEQ PS: Y:	First AA of Sig Pep	Last of Sig Pep	Of AA First Last AA First AA Last AA of ID of of of of AA AA Signal NO: Sig Sig Secreted of AA Pep Pep Portion ORF	Last AA of ORF
87	HCEDO21	97957 03/13/97 209073 05/22/97	97957 Uni-ZAP XR 97 03/13/97 209073 05/22/97	76	1419	1 1419 207	1419	207	207	211	_	20	21	37

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Table 1 summarizes the information corresponding to each "Gene No." described above. The nucleotide sequence identified as "NT SEQ ID NO:X" was assembled from partially homologous ("overlapping") sequences obtained from the "cDNA clone ID" identified in Table 1 and, in some cases, from additional related DNA clones. The overlapping sequences were assembled into a single contiguous sequence of high redundancy (usually three to five overlapping sequences at each nucleotide position), resulting in a final sequence identified as SEQ ID NO:X.

The cDNA Clone ID was deposited on the date and given the corresponding deposit number listed in "ATCC Deposit No:Z and Date." Some of the deposits contain multiple different clones corresponding to the same gene. "Vector" refers to the type of vector contained in the cDNA Clone ID.

"Total NT Seq." refers to the total number of nucleotides in the contig identified by "Gene No." The deposited clone may contain all or most of these sequences, reflected by the nucleotide position indicated as "5' NT of Clone Seq." and the "3' NT of Clone Seq." of SEQ ID NO:X. The nucleotide position of SEQ ID NO:X of the putative start codon (methionine) is identified as "5' NT of Start Codon." Similarly, the nucleotide position of SEQ ID NO:X of the predicted signal sequence is identified as "5' NT of First AA of Signal Pep."

The translated amino acid sequence, beginning with the methionine, is identified as "AA SEQ ID NO:Y," although other reading frames can also be easily translated using known molecular biology techniques. The polypeptides produced by these alternative open reading frames are specifically contemplated by the present invention.

The first and last amino acid position of SEQ ID NO:Y of the predicted signal peptide is identified as "First AA of Sig Pep" and "Last AA of Sig Pep." The predicted first amino acid position of SEQ ID NO:Y of the secreted portion is identified as "Predicted First AA of Secreted Portion." Finally, the amino acid position of SEQ ID NO:Y of the last amino acid in the open reading frame is identified as "Last AA of ORF."

SEQ ID NO:X and the translated SEQ ID NO:Y are sufficiently accurate and otherwise suitable for a variety of uses well known in the art and described further below. For instance, SEQ ID NO:X is useful for designing nucleic acid hybridization probes that will detect nucleic acid sequences contained in SEQ ID NO:X or the cDNA contained in the deposited clone. These probes will also hybridize to nucleic acid molecules in biological samples, thereby enabling a variety of forensic and diagnostic methods of the invention. Similarly, polypeptides identified from SEQ ID NO:Y may be used to generate antibodies which bind specifically to the secreted proteins encoded by the cDNA clones identified in Table 1.

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Nevertheless, DNA sequences generated by sequencing reactions can contain sequencing errors. The errors exist as misidentified nucleotides, or as insertions or deletions of nucleotides in the generated DNA sequence. The erroneously inserted or deleted nucleotides cause frame shifts in the reading frames of the predicted amino acid sequence. In these cases, the predicted amino acid sequence diverges from the actual amino acid sequence, even though the generated DNA sequence may be greater than 99.9% identical to the actual DNA sequence (for example, one base insertion or deletion in an open reading frame of over 1000 bases).

Accordingly, for those applications requiring precision in the nucleotide sequence or the amino acid sequence, the present invention provides not only the generated nucleotide sequence identified as SEQ ID NO:X and the predicted translated amino acid sequence identified as SEQ ID NO:Y, but also a sample of plasmid DNA containing a human cDNA of the invention deposited with the ATCC, as set forth in Table 1. The nucleotide sequence of each deposited clone can readily be determined by sequencing the deposited clone in accordance with known methods. The predicted amino acid sequence can then be verified from such deposits. Moreover, the amino acid sequence of the protein encoded by a particular clone can also be directly determined by peptide sequencing or by expressing the protein in a suitable host cell containing the deposited human cDNA, collecting the protein, and determining its sequence.

The present invention also relates to the genes corresponding to SEQ ID NO:X, SEQ ID NO:Y, or the deposited clone. The corresponding gene can be isolated in accordance with known methods using the sequence information disclosed herein. Such methods include preparing probes or primers from the disclosed sequence and identifying or amplifying the corresponding gene from appropriate sources of genomic material.

Also provided in the present invention are species homologs. Species homologs may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source for the desired homologue.

The polypeptides of the invention can be prepared in any suitable manner. Such polypeptides include isolated naturally occurring polypeptides, recombinantly produced polypeptides, synthetically produced polypeptides, or polypeptides produced by a combination of these methods. Means for preparing such polypeptides are well understood in the art.

The polypeptides may be in the form of the secreted protein, including the mature form, or may be a part of a larger protein, such as a fusion protein (see below).

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It is often advantageous to include an additional amino acid sequence which contains secretory or leader sequences, pro-sequences, sequences which aid in purification . such as multiple histidine residues, or an additional sequence for stability during recombinant production.

The polypeptides of the present invention are preferably provided in an isolated form, and preferably are substantially purified. A recombinantly produced version of a polypeptide, including the secreted polypeptide, can be substantially purified by the one-step method described in Smith and Johnson, Gene 67:31-40 (1988). Polypeptides of the invention also can be purified from natural or recombinant sources using antibodies of the invention raised against the secreted protein in methods which are well known in the art.

Signal Sequences

Methods for predicting whether a protein has a signal sequence, as well as the cleavage point for that sequence, are available. For instance, the method of McGeoch, Virus Res. 3:271-286 (1985), uses the information from a short N-terminal charged region and a subsequent uncharged region of the complete (uncleaved) protein. The method of von Heinje, Nucleic Acids Res. 14:4683-4690 (1986) uses the information from the residues surrounding the cleavage site, typically residues -13 to +2, where +1 indicates the amino terminus of the secreted protein. The accuracy of predicting the cleavage points of known mammalian secretory proteins for each of these methods is in the range of 75-80%. (von Heinje, supra.) However, the two methods do not always produce the same predicted cleavage point(s) for a given protein.

In the present case, the deduced amino acid sequence of the secreted polypeptide was analyzed by a computer program called SignalP (Henrik Nielsen et al., Protein Engineering 10:1-6 (1997)), which predicts the cellular location of a protein based on the amino acid sequence. As part of this computational prediction of localization, the methods of McGeoch and von Heinje are incorporated. The analysis of the amino acid sequences of the secreted proteins described herein by this program provided the results shown in Table 1.

As one of ordinary skill would appreciate, however, cleavage sites sometimes vary from organism to organism and cannot be predicted with absolute certainty. Accordingly, the present invention provides secreted polypeptides having a sequence shown in SEQ ID NO:Y which have an N-terminus beginning within 5 residues (i.e., + or - 5 residues) of the predicted cleavage point. Similarly, it is also recognized that in some cases, cleavage of the signal sequence from a secreted protein is not entirely

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uniform, resulting in more than one secreted species. These polypeptides, and the polynucleotides encoding such polypeptides, are contemplated by the present invention.

Moreover, the signal sequence identified by the above analysis may not necessarily predict the naturally occurring signal sequence. For example, the naturally occurring signal sequence may be further upstream from the predicted signal sequence. However, it is likely that the predicted signal sequence will be capable of directing the secreted protein to the ER. These polypeptides, and the polynucleotides encoding such polypeptides, are contemplated by the present invention.

10 Polynucleotide and Polypeptide Variants

"Variant" refers to a polynucleotide or polypeptide differing from the polynucleotide or polypeptide of the present invention, but retaining essential properties thereof. Generally, variants are overall closely similar, and, in many regions, identical to the polynucleotide or polypeptide of the present invention.

By a polynucleotide having a nucleotide sequence at least, for example, 95% "identical" to a reference nucleotide sequence of the present invention, it is intended that the nucleotide sequence of the polynucleotide is identical to the reference sequence except that the polynucleotide sequence may include up to five point mutations per each 100 nucleotides of the reference nucleotide sequence encoding the polypeptide. In other words, to obtain a polynucleotide having a nucleotide sequence at least 95% identical to a reference nucleotide sequence, up to 5% of the nucleotides in the reference sequence may be deleted or substituted with another nucleotide, or a number of nucleotides up to 5% of the total nucleotides in the reference sequence may be inserted into the reference sequence. The query sequence may be an entire sequence shown in Table 1, the ORF (open reading frame), or any fragement specified as described herein.

As a practical matter, whether any particular nucleic acid molecule or polypeptide is at least 90%, 95%, 96%, 97%, 98% or 99% identical to a nucleotide sequence of the presence invention can be determined conventionally using known computer programs. A preferred method for determing the best overall match between a query sequence (a sequence of the present invention) and a subject sequence, also referred to as a global sequence alignment, can be determined using the FASTDB computer program based on the algorithm of Brutlag et al. (Comp. App. Biosci. (1990) 6:237-245). In a sequence alignment the query and subject sequences are both DNA sequences. An RNA sequence can be compared by converting U's to T's. The result of said global sequence alignment is in percent identity. Preferred parameters used in a FASTDB alignment of DNA sequences to calculate percent identity are:

Matrix=Unitary, k-tuple=4, Mismatch Penalty=1, Joining Penalty=30, Randomization

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Group Length=0, Cutoff Score=1, Gap Penalty=5, Gap Size Penalty 0.05, Window Size=500 or the length of the subject nucleotide sequence, whichever is shorter.

If the subject sequence is shorter than the query sequence because of 5' or 3' deletions, not because of internal deletions, a manual correction must be made to the results. This is because the FASTDB program does not account for 5' and 3' truncations of the subject sequence when calculating percent identity. For subject sequences truncated at the 5' or 3' ends, relative to the the query sequence, the percent identity is corrected by calculating the number of bases of the query sequence that are 5' and 3' of the subject sequence, which are not matched/aligned, as a percent of the total bases of the query sequence. Whether a nucleotide is matched/aligned is determined by results of the FASTDB sequence alignment. This percentage is then subtracted from the percent identity, calculated by the above FASTDB program using the specified parameters, to arrive at a final percent identity score. This corrected score is what is used for the purposes of the present invention. Only bases outside the 5' and 3' bases of the subject sequence, as displayed by the FASTDB alignment, which are not matched/aligned with the query sequence, are calculated for the purposes of manually adjusting the percent identity score.

For example, a 90 base subject sequence is aligned to a 100 base query sequence to determine percent identity. The deletions occur at the 5' end of the subject sequence and therefore, the FASTDB alignment does not show a matched/alignement of the first 10 bases at 5' end. The 10 unpaired bases represent 10% of the sequence (number of bases at the 5' and 3' ends not matched/total number of bases in the query sequence) so 10% is subtracted from the percent identity score calculated by the FASTDB program. If the remaining 90 bases were perfectly matched the final percent identity would be 90%. In another example, a 90 base subject sequence is compared with a 100 base query sequence. This time the deletions are internal deletions so that there are no bases on the 5' or 3' of the subject sequence which are not matched/aligned with the query. In this case the percent identity calculated by FASTDB is not manually corrected. Once again, only bases 5' and 3' of the subject sequence which are not matched/aligned with the query sequence are manually corrected for. No other manual corrections are to made for the purposes of the present invention.

By a polypeptide having an amino acid sequence at least, for example, 95% "identical" to a query amino acid sequence of the present invention, it is intended that the amino acid sequence of the subject polypeptide is identical to the query sequence except that the subject polypeptide sequence may include up to five amino acid alterations per each 100 amino acids of the query amino acid sequence. In other words, to obtain a polypeptide having an amino acid sequence at least 95% identical to a query

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amino acid sequence, up to 5% of the amino acid residues in the subject sequence may be inserted, deleted, (indels) or substituted with another amino acid. These alterations of the reference sequence may occur at the amino or carboxy terminal positions of the reference amino acid sequence or anywhere between those terminal positions, interspersed either individually among residues in the reference sequence or in one or

more contiguous groups within the reference sequence.

As a practical matter, whether any particular polypeptide is at least 90%, 95%, 96%, 97%, 98% or 99% identical to, for instance, the amino acid sequences shown in Table 1 or to the amino acid sequence encoded by deposited DNA clone can be determined conventionally using known computer programs. A preferred method for determing the best overall match between a query sequence (a sequence of the present invention) and a subject sequence, also referred to as a global sequence alignment, can be determined using the FASTDB computer program based on the algorithm of Brutlag et al. (Comp. App. Biosci. (1990) 6:237-245). In a sequence alignment the query and subject sequences are either both nucleotide sequences or both amino acid sequences. The result of said global sequence alignment is in percent identity. Preferred parameters used in a FASTDB amino acid alignment are: Matrix=PAM 0, k-tuple=2, Mismatch Penalty=1, Joining Penalty=20, Randomization Group Length=0, Cutoff Score=1, Window Size=sequence length, Gap Penalty=5, Gap Size Penalty=0.05, Window Size=500 or the length of the subject amino acid sequence, whichever is shorter.

If the subject sequence is shorter than the query sequence due to N- or Cterminal deletions, not because of internal deletions, a manual correction must be made to the results. This is because the FASTDB program does not account for N- and Cterminal truncations of the subject sequence when calculating global percent identity. For subject sequences truncated at the N- and C-termini, relative to the the query sequence, the percent identity is corrected by calculating the number of residues of the query sequence that are N- and C-terminal of the subject sequence, which are not matched/aligned with a corresponding subject residue, as a percent of the total bases of the query sequence. Whether a residue is matched/aligned is determined by results of the FASTDB sequence alignment. This percentage is then subtracted from the percent identity, calculated by the above FASTDB program using the specified parameters, to arrive at a final percent identity score. This final percent identity score is what is used for the purposes of the present invention. Only residues to the N- and C-termini of the subject sequence, which are not matched/aligned with the query sequence, are considered for the purposes of manually adjusting the percent identity score. That is, only query residue positions outside the farthest N- and C-terminal residues of the subject sequence.

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purposes of the present invention.

For example, a 90 amino acid residue subject sequence is aligned with a 100 residue query sequence to determine percent identity. The deletion occurs at the N-terminus of the subject sequence and therefore, the FASTDB alignment does not show a matching/alignment of the first 10 residues at the N-terminus. The 10 unpaired residues represent 10% of the sequence (number of residues at the N- and C-termini not matched/total number of residues in the query sequence) so 10% is subtracted from the percent identity score calculated by the FASTDB program. If the remaining 90 residues were perfectly matched the final percent identity would be 90%. In another example, a 90 residue subject sequence is compared with a 100 residue query sequence. This time the deletions are internal deletions so there are no residues at the N- or C-termini of the subject sequence which are not matched/aligned with the query. In this case the percent identity calculated by FASTDB is not manually corrected. Once again, only residue positions outside the N- and C-terminal ends of the subject sequence, as displayed in the FASTDB alignment, which are not matched/aligned with the query sequence are manually corrected for. No other manual corrections are to made for the

The variants may contain alterations in the coding regions, non-coding regions, or both. Especially preferred are polynucleotide variants containing alterations which produce silent substitutions, additions, or deletions, but do not alter the properties or activities of the encoded polypeptide. Nucleotide variants produced by silent substitutions due to the degeneracy of the genetic code are preferred. Moreover, variants in which 5-10, 1-5, or 1-2 amino acids are substituted, deleted, or added in any combination are also preferred. Polynucleotide variants can be produced for a variety of reasons, e.g., to optimize codon expression for a particular host (change codons in the human mRNA to those preferred by a bacterial host such as E. coli).

Naturally occurring variants are called "allelic variants," and refer to one of several alternate forms of a gene occupying a given locus on a chromosome of an organism. (Genes II, Lewin, B., ed., John Wiley & Sons, New York (1985).) These allelic variants can vary at either the polynucleotide and/or polypeptide level. Alternatively, non-naturally occurring variants may be produced by mutagenesis techniques or by direct synthesis.

Using known methods of protein engineering and recombinant DNA technology, variants may be generated to improve or alter the characteristics of the polypeptides of the present invention. For instance, one or more amino acids can be deleted from the N-terminus or C-terminus of the secreted protein without substantial loss of biological function. The authors of Ron et al., J. Biol. Chem. 268: 2984-2988 (1993), reported variant KGF proteins having heparin binding activity even after

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deleting 3, 8, or 27 amino-terminal amino acid residues. Similarly, Interferon gamma exhibited up to ten times higher activity after deleting 8-10 amino acid residues from the carboxy terminus of this protein. (Dobeli et al., J. Biotechnology 7:199-216 (1988).)

Moreover, ample evidence demonstrates that variants often retain a biological activity similar to that of the naturally occurring protein. For example, Gayle and coworkers (J. Biol. Chem 268:22105-22111 (1993)) conducted extensive mutational analysis of human cytokine IL-1a. They used random mutagenesis to generate over 3,500 individual IL-1a mutants that averaged 2.5 amino acid changes per variant over the entire length of the molecule. Multiple mutations were examined at every possible amino acid position. The investigators found that "[m]ost of the molecule could be altered with little effect on either [binding or biological activity]." (See, Abstract.) In fact, only 23 unique amino acid sequences, out of more than 3,500 nucleotide sequences examined, produced a protein that significantly differed in activity from wild-type.

Furthermore, even if deleting one or more amino acids from the N-terminus or C-terminus of a polypeptide results in modification or loss of one or more biological functions, other biological activities may still be retained. For example, the ability of a deletion variant to induce and/or to bind antibodies which recognize the secreted form will likely be retained when less than the majority of the residues of the secreted form are removed from the N-terminus or C-terminus. Whether a particular polypeptide lacking N- or C-terminal residues of a protein retains such immunogenic activities can readily be determined by routine methods described herein and otherwise known in the art.

Thus, the invention further includes polypeptide variants which show substantial biological activity. Such variants include deletions, insertions, inversions, repeats, and substitutions selected according to general rules known in the art so as have little effect on activity. For example, guidance concerning how to make phenotypically silent amino acid substitutions is provided in Bowie, J. U. et al., Science 247:1306-1310 (1990), wherein the authors indicate that there are two main strategies for studying the tolerance of an amino acid sequence to change.

The first strategy exploits the tolerance of amino acid substitutions by natural selection during the process of evolution. By comparing amino acid sequences in different species, conserved amino acids can be identified. These conserved amino acids are likely important for protein function. In contrast, the amino acid positions where substitutions have been tolerated by natural selection indicates that these positions are not critical for protein function. Thus, positions tolerating amino acid substitution could be modified while still maintaining biological activity of the protein.

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The second strategy uses genetic engineering to introduce amino acid changes at specific positions of a cloned gene to identify regions critical for protein function. For example, site directed mutagenesis or alanine-scanning mutagenesis (introduction of single alanine mutations at every residue in the molecule) can be used. (Cunningham and Wells, Science 244:1081-1085 (1989).) The resulting mutant molecules can then be tested for biological activity.

As the authors state, these two strategies have revealed that proteins are surprisingly tolerant of amino acid substitutions. The authors further indicate which amino acid changes are likely to be permissive at certain amino acid positions in the protein. For example, most buried (within the tertiary structure of the protein) amino acid residues require nonpolar side chains, whereas few features of surface side chains are generally conserved. Moreover, tolerated conservative amino acid substitutions involve replacement of the aliphatic or hydrophobic amino acids Ala, Val, Leu and Ile: replacement of the hydroxyl residues Ser and Thr; replacement of the acidic residues Asp and Glu; replacement of the amide residues Asn and Gln, replacement of the basic residues Lys, Arg, and His; replacement of the aromatic residues Phe, Tyr, and Trp, and replacement of the small-sized amino acids Ala, Ser, Thr, Met, and Gly.

Besides conservative amino acid substitution, variants of the present invention include (i) substitutions with one or more of the non-conserved amino acid residues, where the substituted amino acid residues may or may not be one encoded by the genetic code, or (ii) substitution with one or more of amino acid residues having a substituent group, or (iii) fusion of the mature polypeptide with another compound, such as a compound to increase the stability and/or solubility of the polypeptide (for example, polyethylene glycol), or (iv) fusion of the polypeptide with additional amino acids, such as an IgG Fc fusion region peptide, or leader or secretory sequence, or a sequence facilitating purification. Such variant polypeptides are deemed to be within the scope of those skilled in the art from the teachings herein.

For example, polypeptide variants containing amino acid substitutions of charged amino acids with other charged or neutral amino acids may produce proteins with improved characteristics, such as less aggregation. Aggregation of pharmaceutical formulations both reduces activity and increases clearance due to the aggregate's immunogenic activity. (Pinckard et al., Clin. Exp. Immunol. 2:331-340 (1967); Robbins et al., Diabetes 36: 838-845 (1987); Cleland et al., Crit. Rev. The apeutic Drug Carrier Systems 10:307-377 (1993).)

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Polynucleotide and Polypeptide Fragments

In the present invention, a "polynucleotide fragment" refers to a short polynucleotide having a nucleic acid sequence contained in the deposited clone or shown in SEQ ID NO:X. The short nucleotide fragments are preferably at least about 15 nt, and more preferably at least about 20 nt, still more preferably at least about 30 nt, and even more preferably, at least about 40 nt in length. A fragment "at least 20 nt in length," for example, is intended to include 20 or more contiguous bases from the cDNA sequence contained in the deposited clone or the nucleotide sequence shown in SEQ ID NO:X. These nucleotide fragments are useful as diagnostic probes and primers as discussed herein. Of course, larger fragments (e.g., 50, 150, 500, 600, 2000 nucleotides) are preferred.

Moreover, representative examples of polynucleotide fragments of the invention, include, for example, fragments having a sequence from about nucleotide number 1-50, 51-100, 101-150, 151-200, 201-250, 251-300, 301-350, 351-400, 401-450, 451-500, 501-550, 551-600, 651-700, or 701 to the end of SEQ ID NO:X or the cDNA contained in the deposited clone. In this context "about" includes the particularly recited ranges, larger or smaller by several (5, 4, 3, 2, or 1) nucleotides, at either terminus or at both termini. Preferably, these fragments encode a polypeptide which has biological activity.

In the present invention, a "polypeptide fragment" refers to a short amino acid sequence contained in SEQ ID NO:Y or encoded by the cDNA contained in the deposited clone. Protein fragments may be "free-standing," or comprised within a larger polypeptide of which the fragment forms a part or region, most preferably as a single continuous region. Representative examples of polypeptide fragments of the invention, include, for example, fragments from about amino acid number 1-20, 21-40, 41-60, 61-80, 81-100, 102-120, 121-140, 141-160, or 161 to the end of the coding region. Moreover, polypeptide fragments can be about 20, 30, 40, 50, 60, 70, 80, 90, 100, 110, 120, 130, 140, or 150 amino acids in length. In this context "about" includes the particularly recited ranges, larger or smaller by several (5, 4, 3, 2, or 1) amino acids, at either extreme or at both extremes.

Preferred polypeptide fragments include the secreted protein as well as the mature form. Further preferred polypeptide fragments include the secreted protein or the mature form having a continuous series of deleted residues from the amino or the carboxy terminus, or both. For example, any number of amino acids, ranging from 1-60, can be deleted from the amino terminus of either the secreted polypeptide or the mature form. Similarly, any number of amino acids, ranging from 1-30, can be deleted from the carboxy terminus of the secreted protein or mature form. Furthermore, any

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combination of the above amino and carboxy terminus deletions are preferred. Similarly, polynucleotide fragments encoding these polypeptide fragments are also preferred.

Also preferred are polypeptide and polynucleotide fragments characterized by structural or functional domains, such as fragments that comprise alpha-helix and alpha-helix forming regions, beta-sheet and beta-sheet-forming regions, turn and turn-forming regions, coil and coil-forming regions, hydrophilic regions, hydrophobic regions, alpha amphipathic regions, beta amphipathic regions, flexible regions, surface-forming regions, substrate binding region, and high antigenic index regions. Polypeptide fragments of SEQ ID NO:Y falling within conserved domains are specifically contemplated by the present invention. Moreover, polynucleotide fragments encoding these domains are also contemplated.

Other preferred fragments are biologically active fragments. Biologically active fragments are those exhibiting activity similar, but not necessarily identical, to an activity of the polypeptide of the present invention. The biological-activity of the fragments may include an improved desired activity, or a decreased undesirable activity.

Epitopes & Antibodies

In the present invention, "epitopes" refer to polypeptide fragments having antigenic or immunogenic activity in an animal, especially in a human. A preferred embodiment of the present invention relates to a polypeptide fragment comprising an epitope, as well as the polynucleotide encoding this fragment. A region of a protein molecule to which an antibody can bind is defined as an "antigenic epitope." In contrast, an "immunogenic epitope" is defined as a part of a protein that elicits an antibody response. (See, for instance, Geysen et al., Proc. Natl. Acad. Sci. USA 81:3998- 4002 (1983).)

Fragments which function as epitopes may be produced by any conventional means. (See, e.g., Houghten, R. A., Proc. Natl. Acad. Sci. USA 82:5131-5135 (1985) further described in U.S. Patent No. 4,631,211.)

In the present invention, antigenic epitopes preferably contain a sequence of at least seven, more preferably at least nine, and most preferably between about 15 to about 30 amino acids. Antigenic epitopes are useful to raise antibodies, including monoclonal antibodies, that specifically bind the epitope. (See, for instance, Wilson et al., Cell 37:767-778 (1984); Sutcliffe, J. G. et al., Science 219:660-666 (1983).)

Similarly, immunogenic epitopes can be used to induce antibodies according to methods well known in the art. (See, for instance, Sutcliffe et al., supra; Wilson et al., supra; Chow, M. et al., Proc. Natl. Acad. Sci. USA 82:910-914; and Bittle, F. J. et

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al., J. Gen. Virol. 66:2347-2354 (1985).) A preferred immunogenic epitope includes the secreted protein. The immunogenic epitopes may be presented together with a carrier protein, such as an albumin, to an animal system (such as rabbit or mouse) or, if it is long enough (at least about 25 amino acids), without a carrier. However, immunogenic epitopes comprising as few as 8 to 10 amino acids have been shown to be sufficient to raise antibodies capable of binding to, at the very least, linear epitopes in a denatured polypeptide (e.g., in Western blotting.)

As used herein, the term "antibody" (Ab) or "monoclonal antibody" (Mab) is meant to include intact molecules as well as antibody fragments (such as, for example, Fab and F(ab')2 fragments) which are capable of specifically binding to protein. Fab and F(ab')2 fragments lack the Fc fragment of intact antibody, clear more rapidly from the circulation, and may have less non-specific tissue binding than an intact antibody. (Wahl et al., J. Nucl. Med. 24:316-325 (1983).) Thus, these fragments are preferred, as well as the products of a FAB or other immunoglobulin expression library. Moreover, antibodies of the present invention include chimeric, single chain, and humanized antibodies.

Fusion Proteins

Any polypeptide of the present invention can be used to generate fusion proteins. For example, the polypeptide of the present invention, when fused to a second protein, can be used as an antigenic tag. Antibodies raised against the polypeptide of the present invention can be used to indirectly detect the second protein by binding to the polypeptide. Moreover, because secreted proteins target cellular locations based on trafficking signals, the polypeptides of the present invention can be used as targeting molecules once fused to other proteins.

Examples of domains that can be fused to polypeptides of the present invention include not only heterologous signal sequences, but also other heterologous functional regions. The fusion does not necessarily need to be direct, but may occur through linker sequences.

Moreover, fusion proteins may also be engineered to improve characteristics of the polypeptide of the present invention. For instance, a region of additional amino acids, particularly charged amino acids, may be added to the N-terminus of the polypeptide to improve stability and persistence during purification from the host cell or subsequent handling and storage. Also, peptide moieties may be added to the polypeptide to facilitate purification. Such regions may be removed prior to final preparation of the polypeptide. The addition of peptide moieties to facilitate handling of polypeptides are familiar and routine techniques in the art.

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Moreover, polypeptides of the present invention, including fragments, and specifically epitopes, can be combined with parts of the constant domain of immunoglobulins (IgG), resulting in chimeric polypeptides. These fusion proteins facilitate purification and show an increased half-life in vivo. One reported example describes chimeric proteins consisting of the first two domains of the human CD4-polypeptide and various domains of the constant regions of the heavy or light chains of mammalian immunoglobulins. (EP A 394,827; Traunecker et al., Nature 331:84-86 (1988).) Fusion proteins having disulfide-linked dimeric structures (due to the IgG) can also be more efficient in binding and neutralizing other molecules, than the monomeric secreted protein or protein fragment alone. (Fountoulakis et al., J. Biochem. 270:3958-3964 (1995).)

Similarly, EP-A-O 464 533 (Canadian counterpart 2045869) discloses fusion proteins comprising various portions of constant region of immunoglobulin molecules together with another human protein or part thereof. In many cases, the Fc part in a fusion protein is beneficial in therapy and diagnosis, and thus can result in, for example, improved pharmacokinetic properties. (EP-A 0232 262.) Alternatively, deleting the Fc part after the fusion protein has been expressed, detected, and purified, would be desired. For example, the Fc portion may hinder therapy and diagnosis if the fusion protein is used as an antigen for immunizations. In drug discovery, for example, human proteins, such as hIL-5, have been fused with Fc portions for the purpose of high-throughput screening assays to identify antagonists of hIL-5. (See, D. Bennett et al., J. Molecular Recognition 8:52-58 (1995); K. Johanson et al., J. Biol. Chem. 270:9459-9471 (1995).)

Moreover, the polypeptides of the present invention can be fused to marker sequences, such as a peptide which facilitates purification of the fused polypeptide. In preferred embodiments, the marker amino acid sequence is a hexa-histidine peptide, such as the tag provided in a pQE vector (QIAGEN, Inc., 9259 Eton Avenue, Chatsworth, CA, 91311), among others, many of which are commercially available. As described in Gentz et al., Proc. Natl. Acad. Sci. USA 86:821-824 (1989), for instance, hexa-histidine provides for convenient purification of the fusion protein. Another peptide tag useful for purification, the "HA" tag, corresponds to an epitope derived from the influenza hemagglutinin protein. (Wilson et al., Cell 37:767 (1984).)

Thus, any of these above fusions can be engineered using the polynucleotides or the polypeptides of the present invention.

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Vectors, Host Cells, and Protein Production

The present invention also relates to vectors containing the polynucleotide of the present invention, host cells, and the production of polypeptides by recombinant techniques. The vector may be, for example, a phage, plasmid, viral, or retroviral vector. Retroviral vectors may be replication competent or replication defective. In the latter case, viral propagation generally will occur only in complementing host cells.

The polynucleotides may be joined to a vector containing a selectable marker for propagation in a host. Generally, a plasmid vector is introduced in a precipitate, such as a calcium phosphate precipitate, or in a complex with a charged lipid. If the vector is a virus, it may be packaged in vitro using an appropriate packaging cell line and then transduced into host cells.

The polynucleotide insert should be operatively linked to an appropriate promoter, such as the phage lambda PL promoter, the E. coli lac, trp, phoA and tac promoters, the SV40 early and late promoters and promoters of retroviral LTRs, to name a few. Other suitable promoters will be known to the skilled artisan. The expression constructs will further contain sites for transcription initiation, termination, and, in the transcribed region, a ribosome binding site for translation. The coding portion of the transcripts expressed by the constructs will preferably include a translation initiating codon at the beginning and a termination codon (UAA, UGA or UAG) appropriately positioned at the end of the polypeptide to be translated.

As indicated, the expression vectors will preferably include at least one selectable marker. Such markers include dihydrofolate reductase, G418 or neomycin resistance for eukaryotic cell culture and tetracycline, kanamycin or ampicillin resistance genes for culturing in E. coli and other bacteria. Representative examples of appropriate hosts include, but are not limited to, bacterial cells, such as E. coli, Streptomyces and Salmonella typhimurium cells; fungal cells, such as yeast cells; insect cells such as Drosophila S2 and Spodoptera Sf9 cells; animal cells such as CHO, COS, 293, and Bowes melanoma cells; and plant cells. Appropriate culture mediums and conditions for the above-described host cells are known in the art.

Among vectors preferred for use in bacteria include pQE70, pQE60 and pQE-9, available from QIAGEN, Inc.; pBluescript vectors, Phagescript vectors, pNH8A, pNH16a, pNH18A, pNH46A, available from Stratagene Cloning Systems, Inc.; and ptrc99a, pKK223-3, pKK233-3, pDR540, pRIT5 available from Pharmacia Biotech, Inc. Among preferred eukaryotic vectors are pWLNEO, pSV2CAT, pOG44, pXT1 and pSG available from Stratagene; and pSVK3, pBPV, pMSG and pSVL available from Pharmacia. Other suitable vectors will be readily apparent to the skilled artisan.

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Introduction of the construct into the host cell can be effected by calcium phosphate transfection, DEAE-dextran mediated transfection, cationic lipid-mediated transfection, electroporation, transduction, infection, or other methods. Such methods are described in many standard laboratory manuals, such as Davis et al., Basic Methods In Molecular Biology (1986). It is specifically contemplated that the polypeptides of the present invention may in fact be expressed by a host cell lacking a recombinant vector.

A polypeptide of this invention can be recovered and purified from recombinant cell cultures by well-known methods including ammonium sulfate or ethanol precipitation, acid extraction, anion or cation exchange chromatography, phosphocellulose chromatography, hydrophobic interaction chromatography, affinity chromatography, hydroxylapatite chromatography and lectin chromatography. Most preferably, high performance liquid chromatography ("HPLC") is employed for purification.

Polypeptides of the present invention, and preferably the secreted form, can also be recovered from: products purified from natural sources, including bodily fluids, tissues and cells, whether directly isolated or cultured; products of chemical synthetic procedures; and products produced by recombinant techniques from a prokaryotic or eukaryotic host, including, for example, bacterial, yeast, higher plant, insect, and mammalian cells. Depending upon the host employed in a recombinant production procedure, the polypeptides of the present invention may be glycosylated or may be non-glycosylated. In addition, polypeptides of the invention may also include an initial modified methionine residue, in some cases as a result of host-mediated processes. Thus, it is well known in the art that the N-terminal methionine encoded by the translation initiation codon generally is removed with high efficiency from any protein after translation in all eukaryotic cells. While the N-terminal methionine on most proteins also is efficiently removed in most prokaryotes, for some proteins, this prokaryotic removal process is inefficient, depending on the nature of the amino acid to which the N-terminal methionine is covalently linked.

30 Uses of the Polynucleotides

Each of the polynucleotides identified herein can be used in numerous ways as reagents. The following description should be considered exemplary and utilizes known techniques.

The polynucleotides of the present invention are useful for chromosome identification. There exists an ongoing need to identify new chromosome markers, since few chromosome marking reagents, based on actual sequence data (repeat

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polymorphisms), are presently available. Each polynucleotide of the present invention can be used as a chromosome marker.

Briefly, sequences can be mapped to chromosomes by preparing PCR primers (preferably 15-25 bp) from the sequences shown in SEQ ID NO:X. Primers can be selected using computer analysis so that primers do not span more than one predicted exon in the genomic DNA. These primers are then used for PCR screening of somatic cell hybrids containing individual human chromosomes. Only those hybrids containing the human gene corresponding to the SEQ ID NO:X will yield an amplified fragment.

Similarly, somatic hybrids provide a rapid method of PCR mapping the polynucleotides to particular chromosomes. Three or more clones can be assigned per day using a single thermal cycler. Moreover, sublocalization of the polynucleotides can be achieved with panels of specific chromosome fragments. Other gene mapping strategies that can be used include in situ hybridization, prescreening with labeled flow-sorted chromosomes, and preselection by hybridization to construct chromosome specific-cDNA libraries.

Precise chromosomal location of the polynucleotides can also be achieved using fluorescence in situ hybridization (FISH) of a metaphase chromosomal spread. This technique uses polynucleotides as short as 500 or 600 bases; however, polynucleotides 2,000-4,000 bp are preferred. For a review of this technique, see Verma et al., "Human Chromosomes: a Manual of Basic Techniques," Pergamon Press, New York (1988).

For chromosome mapping, the polynucleotides can be used individually (to mark a single chromosome or a single site on that chromosome) or in panels (for marking multiple sites and/or multiple chromosomes). Preferred polynucleotides correspond to the noncoding regions of the cDNAs because the coding sequences are more likely conserved within gene families, thus increasing the chance of cross hybridization during chromosomal mapping.

Once a polynucleotide has been mapped to a precise chromosomal location, the physical position of the polynucleotide can be used in linkage analysis. Linkage analysis establishes coinheritance between a chromosomal location and presentation of a particular disease. (Disease mapping data are found, for example, in V. McKusick, Mendelian Inheritance in Man (available on line through Johns Hopkins University Welch Medical Library).) Assuming I megabase mapping resolution and one gene per 20 kb, a cDNA precisely localized to a chromosomal region associated with the disease could be one of 50-500 potential causative genes.

Thus, once coinheritance is established, differences in the polynucleotide and the corresponding gene between affected and unaffected individuals can be examined.

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First, visible structural alterations in the chromosomes, such as deletions or translocations, are examined in chromosome spreads or by PCR. If no structural alterations exist, the presence of point mutations are ascertained. Mutations observed in some or all affected individuals, but not in normal individuals, indicates that the mutation may cause the disease. However, complete sequencing of the polypeptide and the corresponding gene from several normal individuals is required to distinguish the mutation from a polymorphism. If a new polymorphism is identified, this polymorphic polypeptide can be used for further linkage analysis.

Furthermore, increased or decreased expression of the gene in affected individuals as compared to unaffected individuals can be assessed using polynucleotides of the present invention. Any of these alterations (altered expression, chromosomal rearrangement, or mutation) can be used as a diagnostic or prognostic marker.

In addition to the foregoing, a polynucleotide can be used to control gene expression through triple helix formation or antisense DNA or RNA. Both methods rely on binding of the polynucleotide to DNA or RNA. For these techniques, preferred polynucleotides are usually 20 to 40 bases in length and complementary to either the region of the gene involved in transcription (triple helix - see Lee et al., Nucl. Acids Res. 6:3073 (1979); Cooney et al., Science 241:456 (1988); and Dervan et al., Science 251:1360 (1991)) or to the mRNA itself (antisense - Okano, J. Neurochem. 56:560 (1991); Oligodeoxy-nucleotides as Antisense Inhibitors of Gene Expression, CRC Press, Boca Raton, FL (1988).) Triple helix formation optimally results in a shut-off of RNA transcription from DNA, while antisense RNA hybridization blocks translation of an mRNA molecule into polypeptide. Both techniques are effective in model systems, and the information disclosed herein can be used to design antisense or triple helix polynucleotides in an effort to treat disease.

Polynucleotides of the present invention are also useful in gene therapy. One goal of gene therapy is to insert a normal gene into an organism having a defective gene, in an effort to correct the genetic defect. The polynucleotides disclosed in the present invention offer a means of targeting such genetic defects in a highly accurate manner. Another goal is to insert a new gene that was not present in the host genome, thereby producing a new trait in the host cell.

The polynucleotides are also useful for identifying individuals from minute biological samples. The United States military, for example, is considering the use of restriction fragment length polymorphism (RFLP) for identification of its personnel. In this technique, an individual's genomic DNA is digested with one or more restriction enzymes, and probed on a Southern blot to yield unique bands for identifying

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personnel. This method does not suffer from the current limitations of "Dog Tags" which can be lost, switched, or stolen, making positive identification difficult. The polynucleotides of the present invention can be used as additional DNA markers for RFLP.

The polynucleotides of the present invention can also be used as an alternative to RFLP, by determining the actual base-by-base DNA sequence of selected portions of an individual's genome. These sequences can be used to prepare PCR primers for amplifying and isolating such selected DNA, which can then be sequenced. Using this technique, individuals can be identified because each individual will have a unique set of DNA sequences. Once an unique ID database is established for an individual, positive identification of that individual, living or dead, can be made from extremely small tissue samples.

Forensic biology also benefits from using DNA-based identification techniques as disclosed herein. DNA sequences taken from very small biological samples such as tissues, e.g., hair or skin, or body fluids, e.g., blood, saliva, semen, etc., can be amplified using PCR. In one prior art technique, gene sequences amplified from polymorphic loci, such as DQa class II HLA gene. are used in forensic biology to identify individuals. (Erlich, H., PCR Technology, Freeman and Co. (1992).) Once these specific polymorphic loci are amplified, they are digested with one or more restriction enzymes, yielding an identifying set of bands on a Southern blot probed with DNA corresponding to the DQa class II HLA gene. Similarly, polynucleotides of the present invention can be used as polymorphic markers for forensic purposes.

There is also a need for reagents capable of identifying the source of a particular tissue. Such need arises, for example, in forensics when presented with tissue of unknown origin. Appropriate reagents can comprise, for example, DNA probes or primers specific to particular tissue prepared from the sequences of the present invention. Panels of such reagents can identify tissue by species and/or by organ type. In a similar fashion, these reagents can be used to screen tissue cultures for contamination.

In the very least, the polynucleotides of the present invention can be used as molecular weight markers on Southern gels, as diagnostic probes for the presence of a specific mRNA in a particular cell type, as a probe to "subtract-out" known sequences in the process of discovering novel polynucleotides, for selecting and making oligomers for attachment to a "gene chip" or other support, to raise anti-DNA antibodies using DNA immunization techniques, and as an antigen to elicit an immune response.

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Uses of the Polypeptides

Each of the polypeptides identified herein can be used in numerous ways. The following description should be considered exemplary and utilizes known techniques.

A polypeptide of the present invention can be used to assay protein levels in a biological sample using antibody-based techniques. For example, protein expression in tissues can be studied with classical immunohistological methods. (Jalkanen, M., et al., J. Cell. Biol. 101:976-985 (1985); Jalkanen, M., et al., J. Cell. Biol. 105:3087-3096 (1987).) Other antibody-based methods useful for detecting protein gene expression include immunoassays, such as the enzyme linked immunosorbent assay (ELISA) and the radioimmunoassay (RIA). Suitable antibody assay labels are known in the art and include enzyme labels, such as, glucose oxidase, and radioisotopes, such as iodine (125I, 121I), carbon (14C), sulfur (35S), tritium (3H), indium (112In). and technetium (99mTc), and fluorescent labels, such as fluorescein and rhodamine, and biotin.

In addition to assaying secreted protein levels in a biological sample, proteins can also be detected in vivo by imaging. Antibody labels or markers for in vivo imaging of protein include those detectable by X-radiography, NMR or ESR. For X-radiography, suitable labels include radioisotopes such as barium or cesium, which emit detectable radiation but are not overtly harmful to the subject. Suitable markers for NMR and ESR include those with a detectable characteristic spin, such as deuterium, which may be incorporated into the antibody by labeling of nutrients for the relevant hybridoma.

A protein-specific antibody or antibody fragment which has been labeled with an appropriate detectable imaging moiety, such as a radioisotope (for example, 131I, 112In, 99mTc), a radio-opaque substance, or a material detectable by nuclear magnetic resonance, is introduced (for example, parenterally, subcutaneously, or intraperitoneally) into the mammal. It will be understood in the art that the size of the subject and the imaging system used will determine the quantity of imaging moiety needed to produce diagnostic images. In the case of a radioisotope moiety, for a human subject, the quantity of radioactivity injected will normally range from about 5 to 20 millicuries of 99mTc. The labeled antibody or antibody fragment will then preferentially accumulate at the location of cells which contain the specific protein. In vivo tumor imaging is described in S.W. Burchiel et al., "Immunopharmaco: inetics of Radiolabeled Antibodies and Their Fragments." (Chapter 13 in Tumor Image g: The Radiochemical Detection of Cancer, S.W. Burchiel and B. A. Rhodes, eds., slasson Publishing Inc. (1982).)

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Thus, the invention provides a diagnostic method of a disorder, which involves (a) assaying the expression of a polypeptide of the present invention in cells or body fluid of an individual; (b) comparing the level of gene expression with a standard gene expression level, whereby an increase or decrease in the assayed polypeptide gene expression level compared to the standard expression level is indicative of a disorder.

Moreover, polypeptides of the present invention can be used to treat disease. For example, patients can be administered a polypeptide of the present invention in an effort to replace absent or decreased levels of the polypeptide (e.g., insulin), to supplement absent or decreased levels of a different polypeptide (e.g., hemoglobin S for hemoglobin B), to inhibit the activity of a polypeptide (e.g., an oncogene), to activate the activity of a polypeptide (e.g., by binding to a receptor), to reduce the activity of a membrane bound receptor by competing with it for free ligand (e.g., soluble TNF receptors used in reducing inflammation), or to bring about a desired response (e.g., blood vessel growth).

Similarly, antibodies directed to a polypeptide of the present invention can also be used to treat disease. For example, administration of an antibody directed to a polypeptide of the present invention can bind and reduce overproduction of the polypeptide. Similarly, administration of an antibody can activate the polypeptide, such as by binding to a polypeptide bound to a membrane (receptor).

At the very least, the polypeptides of the present invention can be used as molecular weight markers on SDS-PAGE gels or on molecular sieve gel filtration columns using methods well known to those of skill in the art. Polypeptides can also be used to raise antibodies, which in turn are used to measure protein expression from a recombinant cell, as a way of assessing transformation of the host cell. Moreover, the polypeptides of the present invention can be used to test the following biological activities.

Biological Activities

The polynucleotides and polypeptides of the present invention can be used in assays to test for one or more biological activities. If these polynucleotides and polypeptides do exhibit activity in a particular assay, it is likely that these molecules may be involved in the diseases associated with the biological activity. Thus, the polynucleotides and polypeptides could be used to treat the associated disease.

35 Immune Activity

A polypeptide or polynucleotide of the present invention may be useful in treating deficiencies or disorders of the immune system, by activating or inhibiting the

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proliferation, differentiation, or mobilization (chemotaxis) of immune cells. Immune cells develop through a process called hematopoiesis, producing myeloid (platelets, red blood cells, neutrophils, and macrophages) and lymphoid (B and T lymphocytes) cells from pluripotent stem cells. The etiology of these immune deficiencies or disorders may be genetic, somatic, such as cancer or some autoimmune disorders, acquired (e.g., by chemotherapy or toxins), or infectious. Moreover, a polynucleotide or polypeptide of the present invention can be used as a marker or detector of a particular immune system disease or disorder.

A polynucleotide or polypeptide of the present invention may be useful in treating or detecting deficiencies or disorders of hematopoietic cells. A polypeptide or polynucleotide of the present invention could be used to increase differentiation and proliferation of hematopoietic cells, including the pluripotent stem cells, in an effort to treat those disorders associated with a decrease in certain (or many) types hematopoietic cells. Examples of immunologic deficiency syndromes include, but are not limited to: blood protein disorders (e.g. agammaglobulinemia, dysgammaglobulinemia), ataxia telangiectasia, common variable immunodeficiency, Digeorge Syndrome, HIV infection, HTLV-BLV infection, leukocyte adhesion deficiency syndrome. lymphopenia, phagocyte bactericidal dysfunction, severe combined immunodeficiency (SCIDs), Wiskott-Aldrich Disorder, anemia, thrombocytopenia, or hemoglobinuria.

Moreover, a polypeptide or polynucleotide of the present invention could also be used to modulate hemostatic (the stopping of bleeding) or thrombolytic activity (clet formation). For example, by increasing hemostatic or thrombolytic activity, a polynucleotide or polypeptide of the present invention could be used to treat blood coagulation disorders (e.g., afibrinogenemia, factor deficiencies), blood platelet disorders (e.g. thrombocytopenia), or wounds resulting from trauma, surgery, or other causes. Alternatively, a polynucleotide or polypeptide of the present invention that can decrease hemostatic or thrombolytic activity could be used to inhibit or dissolve clotting. These molecules could be important in the treatment of heart attacks (infarction), strokes, or scarring.

A polynucleotide or polypeptide of the present invention may also be useful in treating or detecting autoimmune disorders. Many autoimmune disorders result from inappropriate recognition of self as foreign material by immune cells. This inappropriate recognition results in an immune response leading to the destruction of the host tissue. Therefore, the administration of a polypeptide or polynucleotide of the present invention that inhibits an immune response, particularly the proliferation, differentiation, or chemotaxis of T-cells, may be an effective therapy in preventing autoimmune disorders.

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Examples of autoimmune disorders that can be treated or detected by the present invention include, but are not limited to: Addison's Disease, hemolytic anemia, antiphospholipid syndrome, rheumatoid arthritis, dermatitis, allergic encephalomyelitis, glomerulonephritis, Goodpasture's Syndrome, Graves' Disease, Multiple Sclerosis, Myasthenia Gravis, Neuritis, Ophthalmia, Bullous Pemphigoid, Pemphigus, Polyendocrinopathies, Purpura, Reiter's Disease, Stiff-Man Syndrome, Autoimmune Thyroiditis, Systemic Lupus Erythematosus, Autoimmune Pulmonary Inflammation, Guillain-Barre Syndrome, insulin dependent diabetes mellitis, and autoimmune inflammatory eye disease.

Similarly, allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems, may also be treated by a polypeptide or polynucleotide of the present invention. Moreover, these molecules can be used to treat anaphylaxis, hypersensitivity to an antigenic molecule, or blood group incompatibility.

A polynucleotide or polypeptide of the present invention may also be used to treat and/or prevent organ rejection or graft-versus-host disease (GVHD). Organ rejection occurs by host immune cell destruction of the transplanted tissue through an immune response. Similarly, an immune response is also involved in GVHD, but, in this case, the foreign transplanted immune cells destroy the host tissues. The administration of a polypeptide or polynucleotide of the present invention that inhibits an immune response, particularly the proliferation, differentiation, or chemotaxis of T-cells, may be an effective therapy in preventing organ rejection or GVHD.

Similarly, a polypeptide or polynucleotide of the present invention may also be used to modulate inflammation. For example, the polypeptide or polynucleotide may inhibit the proliferation and differentiation of cells involved in an inflammatory response. These molecules can be used to treat inflammatory conditions, both chronic and acute conditions, including inflammation associated with infection (e.g., septic shock, sepsis, or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusion injury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine or chemokine induced lung injury, inflammatory bowel disease, Crohn's disease, or resulting from over production of cytokines (e.g., TNF or IL-1.)

Hyperproliferative Disorders

A polypeptide or polynucleotide can be used to treat or detect hyperproliferative disorders, including neoplasms. A polypeptide or polynucleotide of the present invention may inhibit the proliferation of the disorder through direct or indirect

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interactions. Alternatively, a polypeptide or polynucleotide of the present invention may proliferate other cells which can inhibit the hyperproliferative disorder.

For example, by increasing an immune response, particularly increasing antigenic qualities of the hyperproliferative disorder or by proliferating, differentiating, or mobilizing T-cells, hyperproliferative disorders can be treated. This immune response may be increased by either enhancing an existing immune response, or by initiating a new immune response. Alternatively, decreasing an immune response may also be a method of treating hyperproliferative disorders, such as a chemotherapeutic agent.

Examples of hyperproliferative disorders that can be treated or detected by a polynucleotide or polypeptide of the present invention include, but are not limited to neoplasms located in the: abdomen, bone, breast, digestive system, liver, pancreas, peritoneum, endocrine glands (adrenal, parathyroid, pituitary, testicles, ovary, thymus, thyroid), eye, head and neck, nervous (central and peripheral), lymphatic system, pelvic, skin, soft tissue, spleen, thoracic, and urogenital.

Similarly, other hyperproliferative disorders can also be treated or detected by a polynucleotide or polypeptide of the present invention. Examples of such hyperproliferative disorders include, but are not limited to: hypergammaglobulinemia, lymphoproliferative disorders, paraproteinemias, purpura, sarcoidosis, Sezary Syndrome. Waldenstron's Macroglobulinemia, Gaucher's Disease, histiocytosis, and any other hyperproliferative disease, besides neoplasia, located in an organ system listed above.

Infectious Disease

A polypeptide or polynucleotide of the present invention can be used to treat or detect infectious agents. For example, by increasing the immune response, particularly increasing the proliferation and differentiation of B and/or T cells, infectious diseases may be treated. The immune response may be increased by either enhancing an existing immune response, or by initiating a new immune response. Alternatively, the polypeptide or polynucleotide of the present invention may also directly inhibit the infectious agent, without necessarily eliciting an immune response.

Viruses are one example of an infectious agent that can cause disease or symptoms that can be treated or detected by a polynucleotide or polypeptide of the present invention. Examples of viruses, include, but are not limited to the following DNA and RNA viral families: Arbovirus, Adenoviridae, Arenaviridae, Arterivirus, Birnaviridae, Bunyaviridae, Caliciviridae, Circoviridae, Coronaviridae, Flaviviridae, Hepadnaviridae (Hepatitis). Herpesviridae (such as. Cytomegalovirus. Herpes

Simplex, Herpes Zoster), Mononegavirus (e.g., Paramyxoviridae, Morbillivirus, Rhabdoviridae), Orthomyxoviridae (e.g., Influenza), Papovaviridae, Parvoviridae, Picornaviridae, Poxviridae (such as Smallpox or Vaccinia), Reoviridae (e.g., Rotavirus), Retroviridae (HTLV-I, HTLV-II, Lentivirus), and Togaviridae (e.g., Rubivirus). Viruses falling within these families can cause a variety of diseases or symptoms, including, but not limited to: arthritis, bronchiollitis, encephalitis, eye infections (e.g., conjunctivitis, keratitis), chronic fatigue syndrome, hepatitis (A, B, C, E, Chronic Active, Delta), meningitis, opportunistic infections (e.g., AIDS), pneumonia, Burkitt's Lymphoma, chickenpox, hemorrhagic fever, Measles, Mumps, Parainfluenza, Rabies, the common cold, Polio, leukemia, Rubella, sexually transmitted diseases, skin diseases (e.g., Kaposi's, warts), and viremia. A polypeptide or polynucleotide of the present invention can be used to treat or detect any of these symptoms or diseases.

Similarly, bacterial or fungal agents that can cause disease or symptoms and that can be treated or detected by a polynucleotide or polypeptide of the present invention 15 include, but not limited to, the following Gram-Negative and Gram-positive bacterial families and fungi: Actinomycetales (e.g., Corynebacterium, Mycobacterium, Norcardia), Aspergillosis, Bacillaceae (e.g., Anthrax, Clostridium), Bacteroidaceae, Blastomycosis, Bordetella, Borrelia, Brucellosis, Candidiasis, Campylobacter, Coccidioidomycosis, Cryptococcosis, Dermatocycoses, Enterobacteriaceae (Klebsiella, 20 Salmonella, Serratia, Yersinia), Erysipelothrix, Helicobacter, Legionellosis, Leptospirosis, Listeria, Mycoplasmatales, Neisseriaceae (e.g., Acinetobacter, Gonorrhea, Menigococcal), Pasteurellacea Infections (e.g., Actinobacillus, Heamophilus, Pasteurella), Pseudomonas, Rickettsiaceae, Chlamydiaceae. Syphilis, and Staphylococcal. These bacterial or fungal families can cause the following diseases 25 or symptoms, including, but not limited to: bacteremia, endocarditis, eye infections (conjunctivitis, tuberculosis, uveitis), gingivitis, opportunistic infections (e.g., AIDS related infections), paronychia, prosthesis-related infections, Reiter's Disease, respiratory tract infections, such as Whooping Cough or Empyema, sepsis, Lyme Disease, Cat-Scratch Disease, Dysentery, Paratyphoid Fever, food poisoning, 30 Typhoid, pneumonia, Gonorrhea, meningitis, Chlamydia, Syphilis, Diphtheria, Leprosy, Paratuberculosis, Tuberculosis, Lupus, Botulism, gangrene, tetanus, impetigo, Rheumatic Fever, Scarlet Fever, sexually transmitted diseases, skin diseases (e.g., cellulitis, dermatocycoses), toxemia, urinary tract infections, wound infections. A polypeptide or polynucleotide of the present invention can be used to treat or detect 35 any of these symptoms or diseases.

Moreover, parasitic agents causing disease or symptoms that can be treated or detected by a polynucleotide or polypeptide of the present invention include, but not limited to, the following families: Amebiasis. Babesiosis, Coccidiosis, Cryptosporidiosis, Dientamoebiasis, Dourine, Ectoparasitic, Giardiasis, Helminthiasis, Leishmaniasis, Theileriasis, Toxoplasmosis, Trypanosomiasis, and Trichomonas. These parasites can cause a variety of diseases or symptoms, including, but not limited to: Scabies, Trombiculiasis, eye infections, intestinal disease (e.g., dysentery, giardiasis), liver disease, lung disease, opportunistic infections (e.g., AIDS related), Malaria, pregnancy complications, and toxoplasmosis. A polypeptide or polynucleotide of the present invention can be used to treat or detect any of these symptoms or diseases.

Preferably, treatment using a polypeptide or polynucleotide of the present invention could either be by administering an effective amount of a polypeptide to the patient, or by removing cells from the patient, supplying the cells with a polynucleotide of the present invention, and returning the engineered cells to the patient (ex vivo therapy). Moreover, the polypeptide or polynucleotide of the present invention can be used as an antigen in a vaccine to raise an immune response against infectious disease.

Regeneration

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A polynucleotide or polypeptide of the present invention can be used to differentiate, proliferate, and attract cells, leading to the regeneration of tissues. (See, Science 276:59-87 (1997).) The regeneration of tissues could be used to repair, replace, or protect tissue damaged by congenital defects, trauma (wounds, burns, incisions, or ulcers), age, disease (e.g. osteoporosis, osteocarthritis, periodontal disease, liver failure), surgery, including cosmetic plastic surgery, fibrosis, reperfusion injury, or systemic cytokine damage.

Tissues that could be regenerated using the present invention include organs (e.g., pancreas, liver, intestine, kidney, skin, endothelium), muscle (smooth, skeletal or cardiac), vascular (including vascular endothelium), nervous, hematopoietic, and skeletal (bone, cartilage, tendon, and ligament) tissue. Preferably, regeneration occurs without or decreased scarring. Regeneration also may include angiogenesis.

Moreover, a polynucleotide or polypeptide of the present invention may increase regeneration of tissues difficult to heal. For example, increased tendon/ligament regeneration would quicken recovery time after damage. A polynucleotide or polypeptide of the present invention could also be used prophylactically in an effort to avoid damage. Specific diseases that could be treated include of tendinitis, carpal tunnel syndrome, and other tendon or ligament defects. A further example of tissue

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regeneration of non-healing wounds includes pressure ulcers, ulcers associated with vascular insufficiency, surgical, and traumatic wounds.

Similarly, nerve and brain tissue could also be regenerated by using a polynucleotide or polypeptide of the present invention to proliferate and differentiate nerve cells. Diseases that could be treated using this method include central and peripheral nervous system diseases, neuropathies, or mechanical and traumatic disorders (e.g., spinal cord disorders, head trauma, cerebrovascular disease, and stoke). Specifically, diseases associated with peripheral nerve injuries, peripheral neuropathy (e.g., resulting from chemotherapy or other medical therapies), localized neuropathies, and central nervous system diseases (e.g., Alzheimer's disease, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Drager syndrome), could all be treated using the polynucleotide or polypeptide of the present invention.

15 Chemotaxis

A polynucleotide or polypeptide of the present invention may have chemotaxis activity. A chemotaxic molecule attracts or mobilizes cells (e.g., monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells) to a particular site in the body, such as inflammation, infection, or site of hyperproliferation. The mobilized cells can then fight off and/or heal the particular trauma or abnormality.

A polynucleotide or polypeptide of the present invention may increase chemotaxic activity of particular cells. These chemotactic molecules can then be used to treat inflammation, infection, hyperproliferative disorders, or any immune system disorder by increasing the number of cells targeted to a particular location in the body. For example, chemotaxic molecules can be used to treat wounds and other trauma to tissues by attracting immune cells to the injured location. Chemotactic molecules of the present invention can also attract fibroblasts, which can be used to treat wounds.

It is also contemplated that a polynucleotide or polypeptide of the present invention may inhibit chemotactic activity. These molecules could also be used to treat disorders. Thus, a polynucleotide or polypeptide of the present invention could be used as an inhibitor of chemotaxis.

Binding Activity

A polypeptide of the present invention may be used to screen for molecules that bind to the polypeptide or for molecules to which the polypeptide binds. The binding of the polypeptide and the molecule may activate (agonist), increase, inhibit

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(antagonist), or decrease activity of the polypeptide or the molecule bound. Examples of such molecules include antibodies, oligonucleotides, proteins (e.g., receptors), or small molecules.

Preferably, the molecule is closely related to the natural ligand of the polypeptide, e.g., a fragment of the ligand, or a natural substrate, a ligand, a structural or functional mimetic. (See, Coligan et al., Current Protocols in Immunology 1(2):Chapter 5 (1991).) Similarly, the molecule can be closely related to the natural receptor to which the polypeptide binds, or at least, a fragment of the receptor capable of being bound by the polypeptide (e.g., active site). In either case, the molecule can be rationally designed using known techniques.

Preferably, the screening for these molecules involves producing appropriate cells which express the polypeptide, either as a secreted protein or on the cell membrane. Preferred cells include cells from mammals, yeast, Drosophila, or *E. coli*. Cells expressing the polypeptide (or cell membrane containing the expressed polypeptide) are then preferably contacted with a test compound potentially containing the molecule to observe binding, stimulation, or inhibition of activity of either the polypeptide or the molecule.

The assay may simply test binding of a candidate compound to the polypeptide, wherein binding is detected by a label, or in an assay involving competition with a labeled competitor. Further, the assay may test whether the candidate compound results in a signal generated by binding to the polypeptide.

Alternatively, the assay can be carried out using cell-free preparations, polypeptide/molecule affixed to a solid support, chemical libraries, or natural product mixtures. The assay may also simply comprise the steps of mixing a candidate compound with a solution containing a polypeptide, measuring polypeptide/molecule activity or binding, and comparing the polypeptide/molecule activity or binding to a standard.

Preferably, an ELISA assay can measure polypeptide level or activity in a sample (e.g., biological sample) using a monoclonal or polyclonal antibody. The antibody can measure polypeptide level or activity by either binding, directly or indirectly, to the polypeptide or by competing with the polypeptide for a substrate.

All of these above assays can be used as diagnostic or prognostic markers. The molecules discovered using these assays can be used to treat disease or to bring about a particular result in a patient (e.g., blood vessel growth) by activating or inhibiting the polypeptide/molecule. Moreover, the assays can discover agents which may inhibit or enhance the production of the polypeptide from suitably manipulated cells or tissues.

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Therefore, the invention includes a method of identifying compounds which bind to a polypeptide of the invention comprising the steps of: (a) incubating a candidate binding compound with a polypeptide of the invention; and (b) determining if binding has occurred. Moreover, the invention includes a method of identifying agonists/antagonists comprising the steps of: (a) incubating a candidate compound with a polypeptide of the invention, (b) assaying a biological activity, and (b) determining if a biological activity of the polypeptide has been altered.

Other Activities

A polypeptide or polynucleotide of the present invention may also increase or decrease the differentiation or proliferation of embryonic stem cells, besides, as discussed above, hematopoietic lineage.

A polypeptide or polynucleotide of the present invention may also be used to modulate mammalian characteristics, such as body height, weight, hair color, eye color, skin, percentage of adipose tissue, pigmentation, size, and shape (e.g., cosmetic surgery). Similarly, a polypeptide or polynucleotide of the present invention may be used to modulate mammalian metabolism affecting catabolism, anabolism, processing, utilization, and storage of energy.

A polypeptide or polynucleotide of the present invention may be used to change a mammal's mental state or physical state by influencing biorhythms, caricadic rhythms, depression (including depressive disorders), tendency for violence, tolerance for pain, reproductive capabilities (preferably by Activin or Inhibin-like activity), hormonal or endocrine levels, appetite, libido, memory, stress, or other cognitive qualities.

A polypeptide or polynucleotide of the present invention may also be used as a food additive or preservative, such as to increase or decrease storage capabilities, fat content, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional components.

30 Other Preferred Embodiments

Other preferred embodiments of the claimed invention include an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 50 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1.

Also preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of

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positions beginning with the nucleotide at about the position of the 5' Nucleotide of the Clone Sequence and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Also preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of positions beginning with the nucleotide at about the position of the 5' Nucleotide of the Start Codon and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Similarly preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of positions beginning with the nucleotide at about the position of the 5' Nucleotide of the First Amino Acid of the Signal Peptide and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 150 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X.

Further preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 500 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X.

A further preferred embodiment is a nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the nucleotide sequence of SEQ ID NO:X beginning with the nucleotide at about the position of the 5' Nucleotide of the First Amino Acid of the Signal Peptide and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the complete nucleotide sequence of SEQ ID NO:X.

Also preferred is an isolated nucleic acid molecule which hybridizes under stringent hybridization conditions to a nucleic acid molecule, wherein said nucleic acid molecule which hybridizes does not hybridize under stringent hybridization conditions to a nucleic acid molecule having a nucleotide sequence consisting of only A residues or of only T residues.

Also preferred is a composition of matter comprising a DNA molecule which comprises a human cDNA clone identified by a cDNA Clone Identifier in Table 1, which DNA molecule is contained in the material deposited with the American Type

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Culture Collection and given the ATCC Deposit Number shown in Table 1 for said cDNA Clone Identifier.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least 50 contiguous nucleotides in the nucleotide sequence of a human cDNA clone identified by a cDNA Clone Identifier in Table 1, which DNA molecule is contained in the deposit given the ATCC Deposit Number shown in Table 1.

Also preferred is an isolated nucleic acid molecule, wherein said sequence of at least 50 contiguous nucleotides is included in the nucleotide sequence of the complete open reading frame sequence encoded by said human cDNA clone.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to sequence of at least 150 contiguous nucleotides in the nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to sequence of at least 500 contiguous nucleotides in the nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the complete nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is a method for detecting in a biological sample a nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1; which method comprises a step of comparing a nucleotide sequence of at least one nucleic acid molecule in said sample with a sequence selected from said group and determining whether the sequence of said nucleic acid molecule in said sample is at least 95% identical to said selected sequence.

Also preferred is the above method wherein said step of comparing sequences comprises determining the extent of nucleic acid hybridization between nucleic acid molecules in said sample and a nucleic acid molecule comprising said sequence selected from said group. Similarly, also preferred is the above method wherein said step of comparing sequences is performed by comparing the nucleotide sequence determined from a nucleic acid molecule in said sample with said sequence selected from said group. The nucleic acid molecules can comprise DNA molecules or RNA molecules.

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A further preferred embodiment is a method for identifying the species, tissue or cell type of a biological sample which method comprises a step of detecting nucleic acid molecules in said sample, if any, comprising a nucleotide sequence that is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

The method for identifying the species, tissue or cell type of a biological sample can comprise a step of detecting nucleic acid molecules comprising a nucleotide sequence in a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from said group.

Also preferred is a method for diagnosing in a subject a pathological condition associated with abnormal structure or expression of a gene encoding a secreted protein identified in Table 1, which method comprises a step of detecting in a biological sample obtained from said subject nucleic acid molecules, if any, comprising a nucleotide sequence that is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

The method for diagnosing a pathological condition can comprise a step of detecting nucleic acid molecules comprising a nucleotide sequence in a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from said group.

Also preferred is a composition of matter comprising isolated nucleic acid molecules wherein the nucleotide sequences of said nucleic acid molecules comprise a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1. The nucleic acid molecules can comprise DNA molecules or RNA molecules.

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Also preferred is an isolated polypeptide comprising an amino acid sequence at least 90% identical to a sequence of at least about 10 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1.

Also preferred is a polypeptide, wherein said sequence of contiguous amino acids is included in the amino acid sequence of SEQ ID NO:Y in the range of positions beginning with the residue at about the position of the First Amino Acid of the Secreted Portion and ending with the residue at about the Last Amino Acid of the Open Reading Frame as set forth for SEQ ID NO:Y in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 30 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 100 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to the complete amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 90% identical to a sequence of at least about 10 contiguous amino acids in the complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is a polypeptide wherein said sequence of contiguous amino acids is included in the amino acid sequence of a secreted portion of the secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 30 contiguous amino acids in the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 100 contiguous amino acids in the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

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Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is an isolated antibody which binds specifically to a polypeptide comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is a method for detecting in a biological sample a polypeptide comprising an amino acid sequence which is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1: and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1; which method comprises a step of comparing an amino acid sequence of at least one polypeptide molecule in said sample with a sequence selected from said group and determining whether the sequence of said polypeptide molecule in said sample is at least 90% identical to said sequence of at least 10 contiguous amino acids.

Also preferred is the above method wherein said step of comparing an amino acid sequence of at least one polypeptide molecule in said sample with a sequence selected from said group comprises determining the extent of specific binding of polypeptides in said sample to an antibody which binds specifically to a polypeptide comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is the above method wherein said step of comparing sequences is performed by comparing the amino acid sequence determined from a polypeptide molecule in said sample with said sequence selected from said group.

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Also preferred is a method for identifying the species, tissue or cell type of a biological sample which method comprises a step of detecting polypeptide molecules in said sample, if any, comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is the above method for identifying the species, tissue or cell type of a biological sample, which method comprises a step of detecting polypeptide molecules comprising an amino acid sequence in a panel of at least two amino acid sequences, wherein at least one sequence in said panel is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the above group.

Also preferred is a method for diagnosing in a subject a pathological condition associated with abnormal structure or expression of a gene encoding a secreted protein identified in Table 1, which method comprises a step of detecting in a biological sample obtained from said subject polypeptide molecules comprising an amino acid sequence in a panel of at least two amino acid sequences, wherein at least one sequence in said panel is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

In any of these methods, the step of detecting said polypeptide molecules includes using an antibody.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a nucleotide sequence encoding a polypeptide wherein said polypeptide comprises an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

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Also preferred is an isolated nucleic acid molecule, wherein said nucleotide sequence encoding a polypeptide has been optimized for expression of said polypeptide in a prokaryotic host.

Also preferred is an isolated nucleic acid molecule, wherein said polypeptide comprises an amino acid sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is a method of making a recombinant vector comprising inserting any of the above isolated nucleic acid molecule into a vector. Also preferred is the recombinant vector produced by this method. Also preferred is a method of making a recombinant host cell comprising introducing the vector into a host cell, as well as the recombinant host cell produced by this method.

Also preferred is a method of making an isolated polypeptide comprising. culturing this recombinant host cell under conditions such that said polypeptide is expressed and recovering said polypeptide. Also preferred is this method of making an isolated polypeptide, wherein said recombinant host cell is a eukaryotic cell and said polypeptide is a secreted portion of a human secreted protein comprising an amino acid sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y beginning with the residue at the position of the First Amino Acid of the Secreted Portion of SEQ ID NO:Y wherein Y is an integer set forth in Table 1 and said position of the First Amino Acid of the Secreted Portion of SEQ ID NO:Y is defined in Table 1; and an amino acid sequence of a secreted portion of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1. The isolated polypeptide produced by this method is also preferred.

Also preferred is a method of treatment of an individual in need of an increased level of a secreted protein activity, which method comprises administering to such an individual a pharmaceutical composition comprising an amount of an isolated polypeptide, polynucleotide, or antibody of the claimed invention effective to increase the level of said protein activity in said individual.

Having generally described the invention, the same will be more readily understood by reference to the following examples, which are provided by way of illustration and are not intended as limiting.

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Examples

Example 1: Isolation of a Selected cDNA Clone From the Deposited Sample

Each cDNA clone in a cited ATCC deposit is contained in a plasmid vector. Table 1 identifies the vectors used to construct the cDNA library from which each clone was isolated. In many cases, the vector used to construct the library is a phage vector from which a plasmid has been excised. The table immediately below correlates the related plasmid for each phage vector used in constructing the cDNA library. For example, where a particular clone is identified in Table 1 as being isolated in the vector "Lambda Zap." the corresponding deposited clone is in "pBluescript."

	Vector Used to Construct Library	Corresponding Deposited Plasmid
	Lambda Zap	pBluescript (pBS)
	Uni-Zap XR	pBluescript (pBS)
15	Zap Express	pBK
	lafmid BA	plafmid BA
	pSport1	pSport1
	pCMVSport 2.0	pCMVSport 2.0
	pCMVSport 3.0	pCMVSport 3.0
20	pCR®2.1	pCR [®] 2.1

Vectors Lambda Zap (U.S. Patent Nos. 5,128,256 and 5,286,636), Uni-Zap XR (U.S. Patent Nos. 5,128, 256 and 5,286,636), Zap Express (U.S. Patent Nos. 5,128,256 and 5,286,636), pBluescript (pBS) (Short, J. M. et al., Nucleic Acids Res. 16:7583-7600 (1988); Alting-Mees, M. A. and Short, J. M., Nucleic Acids Res.

- 17:9494 (1989)) and pBK (Alting-Mees, M. A. et al., Strategies 5:58-61 (1992)) are commercially available from Stratagene Cloning Systems, Inc., 11011 N. Torrey Pines Road, La Jolla, CA, 92037. pBS contains an ampicillin resistance gene and pBK contains a neomycin resistance gene. Both can be transformed into E. coli strain XL-1 Blue, also available from Stratagene. pBS comes in 4 forms SK+, SK-, KS+ and KŚ.
- The S and K refers to the orientation of the polylinker to the T7 and T3 primer sequences which flank the polylinker region ("S" is for SacI and "K" is for KpnI which are the first sites on each respective end of the linker). "+" or "-" refer to the orientation of the f1 origin of replication ("ori"), such that in one orientation, single stranded rescue initiated from the f1 ori generates sense strand DNA and in the other, antisense.

Vectors pSport1, pCMVSport 2.0 and pCMVSport 3.0, were obtained from Life Technologies, Inc., P. O. Box 6009, Gaithersburg, MD 20897. All Sport vectors contain an ampicillin resistance gene and may be transformed into E. coli strain

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DH10B, also available from Life Technologies. (See, for instance, Gruber, C. E., et al., Focus 15:59 (1993).) Vector lafmid BA (Bento Soares, Columbia University, NY) contains an ampicillin resistance gene and can be transformed into E. coli strain XL-1 Blue. Vector pCR®2.1, which is available from Invitrogen, 1600 Faraday Avenue,

Carlsbad, CA 92008, contains an ampicillin resistance gene and may be transformed into E. coli strain DH10B, available from Life Technologies. (See, for instance, Clark, J. M., Nuc. Acids Res. 16:9677-9686 (1988) and Mead, D. et al., Bio/Technology 9: (1991).) Preferably, a polynucleotide of the present invention does not comprise the phage vector sequences identified for the particular clone in Table 1, as well as the corresponding plasmid vector sequences designated above.

The deposited material in the sample assigned the ATCC Deposit Number cited in Table 1 for any given cDNA clone also may contain one or more additional plasmids, each comprising a cDNA clone different from that given clone. Thus, deposits sharing the same ATCC Deposit Number contain at least a plasmid for each cDNA clone identified in Table 1. Typically, each ATCC deposit sample cited in Table 1 comprises a mixture of approximately equal amounts (by weight) of about 50 plasmid DNAs, each containing a different cDNA clone; but such a deposit sample may include plasmids for more or less than 50 cDNA clones, up to about 500 cDNA clones.

Two approaches can be used to isolate a particular clone from the deposited sample of plasmid DNAs cited for that clone in Table 1. First, a plasmid is directly isolated by screening the clones using a polynucleotide probe corresponding to SEQ ID NO:X.

Particularly, a specific polynucleotide with 30-40 nucleotides is synthesized using an Applied Biosystems DNA synthesizer according to the sequence reported. The oligonucleotide is labeled, for instance, with ³²P-γ-ATP using T4 polynucleotide 25 kinase and purified according to routine methods. (E.g., Maniatis et al., Molecular Cloning: A Laboratory Manual, Cold Spring Harbor Press, Cold Spring, NY (1982).) The plasmid mixture is transformed into a suitable host, as indicated above (such as XL-1 Blue (Stratagene)) using techniques known to those of skill in the art, such as those provided by the vector supplier or in related publications or patents cited above. 30 The transformants are plated on 1.5% agar plates (containing the appropriate selection agent, e.g., ampicillin) to a density of about 150 transformants (colonies) per plate. These plates are screened using Nylon membranes according to routine methods for bacterial colony screening (e.g., Sambrook et al., Molecular Cloning: A Laboratory Manual, 2nd Edit., (1989), Cold Spring Harbor Laboratory Press, pages 1.93 to 35 1.104), or other techniques known to those of skill in the art.

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Alternatively, two primers of 17-20 nucleotides derived from both ends of the SEQ ID NO:X (i.e., within the region of SEQ ID NO:X bounded by the 5' NT and the 3' NT of the clone defined in Table 1) are synthesized and used to amplify the desired cDNA using the deposited cDNA plasmid as a template. The polymerase chain reaction is carried out under routine conditions, for instance, in 25 µl of reaction mixture with 0.5 ug of the above cDNA template. A convenient reaction mixture is 1.5-5 mM MgCl₂, 0.01% (w/v) gelatin, 20 µM each of dATP, dCTP, dGTP, dTTP, 25 pmol of each primer and 0.25 Unit of Taq polymerase. Thirty five cycles of PCR (denaturation at 94°C for 1 min; annealing at 55°C for 1 min; elongation at 72°C for 1 min) are performed with a Perkin-Elmer Cetus automated thermal cycler. The amplified product is analyzed by agarose gel electrophoresis and the DNA band with expected molecular weight is excised and purified. The PCR product is verified to be the selected sequence by subcloning and sequencing the DNA product.

Several methods are available for the identification of the 5' or 3' non-coding portions of a gene which may not be present in the deposited clone. These methods include but are not limited to, filter probing, clone enrichment using specific probes, and protocols similar or identical to 5' and 3' "RACE" protocols which are well known in the art. For instance, a method similar to 5' RACE is available for generating the missing 5' end of a desired full-length transcript. (Fromont-Racine et al., Nucleic Acids Res. 21(7):1683-1684 (1993).)

Briefly, a specific RNA oligonucleotide is ligated to the 5' ends of a population of RNA presumably containing full-length gene RNA transcripts. A primer set containing a primer specific to the ligated RNA oligonucleotide and a primer specific to a known sequence of the gene of interest is used to PCR amplify the 5' portion of the desired full-length gene. This amplified product may then be sequenced and used to generate the full length gene.

This above method starts with total RNA isolated from the desired source, although poly-A+ RNA can be used. The RNA preparation can then be treated with phosphatase if necessary to eliminate 5' phosphate groups on degraded or damaged RNA which may interfere with the later RNA ligase step. The phosphatase should then be inactivated and the RNA treated with tobacco acid pyrophosphatase in order to remove the cap structure present at the 5' ends of messenger RNAs. This reaction leaves a 5' phosphate group at the 5' end of the cap cleaved RNA which can then be ligated to an RNA oligonucleotide using T4 RNA ligase.

This modified RNA preparation is used as a template for first strand cDNA synthesis using a gene specific oligonucleotide. The first strand synthesis reaction is

used as a template for PCR amplification of the desired 5' end using a primer specific to the ligated RNA oligonucleotide and a primer specific to the known sequence of the gene of interest. The resultant product is then sequenced and analyzed to confirm that the 5' end sequence belongs to the desired gene.

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Example 2: Isolation of Genomic Clones Corresponding to a Polynucleotide

A human genomic P1 library (Genomic Systems, Inc.) is screened by PCR using primers selected for the cDNA sequence corresponding to SEQ ID NO:X., according to the method described in Example 1. (See also, Sambrook.)

Example 3: Tissue Distribution of Polypeptide

Tissue distribution of mRNA expression of polynucleotides of the present invention is determined using protocols for Northern blot analysis, described by, among others, Sambrook et al. For example, a cDNA probe produced by the method described in Example 1 is labeled with P³² using the rediprimeTM DNA labeling system (Amersham Life Science), according to manufacturer's instructions. After labeling, the probe is purified using CHROMA SPIN-100TM column (Clontech Laboratories, Inc.), according to manufacturer's protocol number PT1200-1. The purified labeled probe is then used to examine various human tissues for mRNA expression.

Multiple Tissue Northern (MTN) blots containing various human tissues (H) or human immune system tissues (IM) (Clontech) are examined with the labeled probe using ExpressHybTM hybridization solution (Clontech) according to manufacturer's protocol number PT1190-1. Following hybridization and washing, the blots are mounted and exposed to film at -70°C overnight, and the films developed according to standard procedures.

Example 4: Chromosomal Mapping of the Polynucleotides

An oligonucleotide primer set is designed according to the sequence at the 5' end of SEQ ID NO:X. This primer preferably spans about 100 nucleotides. This primer set is then used in a polymerase chain reaction under the following set of conditions: 30 seconds, 95°C; 1 minute, 56°C; 1 minute, 70°C. This cycle is repeated 32 times followed by one 5 minute cycle at 70°C. Human, mouse, and hamster DNA is used as template in addition to a somatic cell hybrid panel containing individual chromosomes or chromosome fragments (Bios, Inc). The reactions is analyzed on

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either 8% polyacrylamide gels or 3.5 % agarose gels. Chromosome mapping is determined by the presence of an approximately 100 bp PCR fragment in the particular somatic cell hybrid.

5 Example 5: Bacterial Expression of a Polypeptide

A polynucleotide encoding a polypeptide of the present invention is amplified using PCR oligonucleotide primers corresponding to the 5' and 3' ends of the DNA sequence, as outlined in Example 1, to synthesize insertion fragments. The primers used to amplify the cDNA insert should preferably contain restriction sites, such as BamHI and XbaI, at the 5' end of the primers in order to clone the amplified product into the expression vector. For example, BamHI and XbaI correspond to the restriction enzyme sites on the bacterial expression vector pQE-9. (Qiagen, Inc., Chatsworth, CA). This plasmid vector encodes antibiotic resistance (Amp^r), a bacterial origin of replication (ori), an IPTG-regulatable promoter/operator (P/O), a ribosome binding site (RBS), a 6-histidine tag (6-His), and restriction enzyme cloning sites.

The pQE-9 vector is digested with BamHI and XbaI and the amplified fragment is ligated into the pQE-9 vector maintaining the reading frame initiated at the bacterial RBS. The ligation mixture is then used to transform the E. coli strain M15/rep4 (Qiagen, Inc.) which contains multiple copies of the plasmid pREP4, which expresses the lacI repressor and also confers kanamycin resistance (Kan^r). Transformants are identified by their ability to grow on LB plates and ampicillin/kanamycin resistant colonies are selected. Plasmid DNA is isolated and confirmed by restriction analysis.

Clones containing the desired constructs are grown overnight (O/N) in liquid culture in LB media supplemented with both Amp (100 ug/ml) and Kan (25 ug/ml). The O/N culture is used to inoculate a large culture at a ratio of 1:100 to 1:250. The cells are grown to an optical density 600 (O.D.⁶⁰⁰) of between 0.4 and 0.6. IPTG (Isopropyl-B-D-thiogalacto pyranoside) is then added to a final concentration of 1 mM. IPTG induces by inactivating the lacI repressor, clearing the P/O leading to increased gene expression.

Cells are grown for an extra 3 to 4 hours. Cells are then harvested by centrifugation (20 mins at 6000Xg). The cell pellet is solubilized in the chaotropic agent 6 Molar Guanidine HCl by stirring for 3-4 hours at 4°C. The cell debris is removed by centrifugation, and the supernatant containing the polypeptide is loaded onto a nickel-nitrilo-tri-acetic acid ("Ni-NTA") affinity resin column (available from QIAGEN, Inc., supra). Proteins with a 6 x His tag bind to the Ni-NTA resin with high

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affinity and can be purified in a simple one-step procedure (for details see: The QIAexpressionist (1995) QIAGEN, Inc., supra).

Briefly, the supernatant is loaded onto the column in 6 M guanidine-HCl, pH 8, the column is first washed with 10 volumes of 6 M guanidine-HCl, pH 8, then washed with 10 volumes of 6 M guanidine-HCl pH 6, and finally the polypeptide is eluted with 6 M guanidine-HCl, pH 5.

The purified protein is then renatured by dialyzing it against phosphate-buffered saline (PBS) or 50 mM Na-acetate, pH 6 buffer plus 200 mM NaCl. Alternatively, the protein can be successfully refolded while immobilized on the Ni-NTA column. The recommended conditions are as follows: renature using a linear 6M-1M urea gradient in 500 mM NaCl, 20% glycerol, 20 mM Tris/HCl pH 7.4, containing protease inhibitors. The renaturation should be performed over a period of 1.5 hours or more. After renaturation the proteins are eluted by the addition of 250 mM immidazole. Immidazole is removed by a final dialyzing step against PBS or 50 mM sodium acetate pH 6 buffer plus 200 mM NaCl. The purified protein is stored at 4°C or frozen at -80°C.

In addition to the above expression vector, the present invention further includes an expression vector comprising phage operator and promoter elements operatively linked to a polynucleotide of the present invention, called pHE4a. (ATCC Accession Number 209645, deposited on February 25, 1998.) This vector contains: 1) a neomycinphosphotransferase gene as a selection marker, 2) an E. coli origin of replication, 3) a T5 phage promoter sequence, 4) two lac operator sequences, 5) a Shine-Delgarno sequence, and 6) the lactose operon repressor gene (lacIq). The origin of replication (oriC) is derived from pUC19 (LTI, Gaithersburg, MD). The promoter sequence and operator sequences are made synthetically.

DNA can be inserted into the pHEa by restricting the vector with NdeI and XbaI, BamHI, XhoI, or Asp718, running the restricted product on a gel, and isolating the larger fragment (the stuffer fragment should be about 310 base pairs). The DNA insert is generated according to the PCR protocol described in Example 1, using PCR primers having restriction sites for NdeI (5' primer) and XbaI, BamHI, XhoI, or Asp718 (3' primer). The PCR insert is gel purified and restricted with compatible enzymes. The insert and vector are ligated according to standard protocols.

The engineered vector could easily be substituted in the above protocol to express protein in a bacterial system.

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Example 6: Purification of a Polypeptide from an Inclusion Body

The following alternative method can be used to purify a polypeptide expressed in *E coli* when it is present in the form of inclusion bodies. Unless otherwise specified, all of the following steps are conducted at 4-10°C.

Upon completion of the production phase of the *E. coli* fermentation, the cell culture is cooled to 4-10°C and the cells harvested by continuous centrifugation at 15,000 rpm (Heraeus Sepatech). On the basis of the expected yield of protein per unit weight of cell paste and the amount of purified protein required, an appropriate amount of cell paste, by weight, is suspended in a buffer solution containing 100 mM Tris, 50 mM EDTA, pH 7.4. The cells are dispersed to a homogeneous suspension using a high shear mixer.

The cells are then lysed by passing the solution through a microfluidizer (Microfuidics, Corp. or APV Gaulin, Inc.) twice at 4000-6000 psi. The homogenate is then mixed with NaCl solution to a final concentration of 0.5 M NaCl, followed by centrifugation at 7000 xg for 15 min. The resultant pellet is washed again using 0.5M NaCl, 100 mM Tris, 50 mM EDTA, pH 7.4.

The resulting washed inclusion bodies are solubilized with 1.5 M guanidine hydrochloride (GuHCl) for 2-4 hours. After 7000 xg centrifugation for 15 min., the pellet is discarded and the polypeptide containing supernatant is incubated at 4°C overnight to allow further GuHCl extraction.

Following high speed centrifugation (30,000 xg) to remove insoluble particles, the GuHCl solubilized protein is refolded by quickly mixing the GuHCl extract with 20 volumes of buffer containing 50 mM sodium, pH 4.5, 150 mM NaCl, 2 mM EDTA by vigorous stirring. The refolded diluted protein solution is kept at 4°C without mixing for 12 hours prior to further purification steps.

To clarify the refolded polypeptide solution, a previously prepared tangential filtration unit equipped with 0.16 µm membrane filter with appropriate surface area (e.g., Filtron), equilibrated with 40 mM sodium acetate, pH 6.0 is employed. The filtered sample is loaded onto a cation exchange resin (e.g., Poros HS-50, Perseptive Biosystems). The column is washed with 40 mM sodium acetate, pH 6.0 and eluted with 250 mM, 500 mM, 1000 mM, and 1500 mM NaCl in the same buffer, in a stepwise manner. The absorbance at 280 nm of the effluent is continuously monitored. Fractions are collected and further analyzed by SDS-PAGE.

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Fractions containing the polypeptide are then pooled and mixed with 4 volumes of water. The diluted sample is then loaded onto a previously prepared set of tandem columns of strong anion (Poros HQ-50, Perseptive Biosystems) and weak anion (Poros CM-20, Perseptive Biosystems) exchange resins. The columns are equilibrated with 40 mM sodium acetate, pH 6.0. Both columns are washed with 40 mM sodium acetate, pH 6.0, 200 mM NaCl. The CM-20 column is then eluted using a 10 column volume linear gradient ranging from 0.2 M NaCl, 50 mM sodium acetate, pH 6.0 to 1.0 M NaCl, 50 mM sodium acetate, pH 6.5. Fractions are collected under constant A₂₈₀ monitoring of the effluent. Fractions containing the polypeptide (determined, for instance, by 16% SDS-PAGE) are then pooled.

The resultant polypeptide should exhibit greater than 95% purity after the above refolding and purification steps. No major contaminant bands should be observed from Commassie blue stained 16% SDS-PAGE gel when 5 μ g of purified protein is loaded. The purified protein can also be tested for endotoxin/LPS contamination, and typically the LPS content is less than 0.1 ng/ml according to LAL assays.

Example 7: Cloning and Expression of a Polypeptide in a Baculovirus Expression System

In this example, the plasmid shuttle vector pA2 is used to insert a polynucleotide into a baculovirus to express a polypeptide. This expression vector contains the strong polyhedrin promoter of the *Autographa californica* nuclear polyhedrosis virus (AcMNPV) followed by convenient restriction sites such as BamHI, Xba I and Asp718. The polyadenylation site of the simian virus 40 ("SV40") is used for efficient polyadenylation. For easy selection of recombinant virus, the plasmid contains the beta-galactosidase gene from *E. coli* under control of a weak Drosophila promoter in the same orientation, followed by the polyadenylation signal of the polyhedrin gene. The inserted genes are flanked on both sides by viral sequences for cell-mediated homologous recombination with wild-type viral DNA to generate a viable virus that express the cloned polynucleotide.

Many other baculovirus vectors can be used in place of the vector above, such as pAc373, pVL941, and pAcIM1, as one skilled in the art would readily appreciate, as long as the construct provides appropriately located signals for transcription, translation, secretion and the like, including a signal peptide and an in-frame AUG as required. Such vectors are described, for instance, in Luckow et al., Virology 170:31-39 (1989).

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Specifically, the cDNA sequence contained in the deposited clone, including the AUG initiation codon and the naturally associated leader sequence identified in Table 1, is amplified using the PCR protocol described in Example 1. If the naturally occurring signal sequence is used to produce the secreted protein, the pA2 vector does not need a second signal peptide. Alternatively, the vector can be modified (pA2 GP) to include a baculovirus leader sequence, using the standard methods described in Summers et al., "A Manual of Methods for Baculovirus Vectors and Insect Cell Culture Procedures," Texas Agricultural Experimental Station Bulletin No. 1555 (1987).

The amplified fragment is isolated from a 1% agarose gel using a commercially available kit ("Geneclean," BIO 101 Inc., La Jolla, Ca.). The fragment then is digested with appropriate restriction enzymes and again purified on a 1% agarose gel.

The plasmid is digested with the corresponding restriction enzymes and optionally, can be dephosphorylated using calf intestinal phosphatase, using routine procedures known in the art. The DNA is then isolated from a 1% agarose gel using a commercially available kit ("Geneclean" BIO 101 Inc., La Jolla, Ca.).

The fragment and the dephosphorylated plasmid are ligated together with T4 DNA ligase. E. coli HB101 or other suitable E. coli hosts such as XL-1 Blue (Stratagene Cloning Systems, La Jolla, CA) cells are transformed with the ligation mixture and spread on culture plates. Bacteria containing the plasmid are identified by digesting DNA from individual colonies and analyzing the digestion product by gel electrophoresis. The sequence of the cloned fragment is confirmed by DNA sequencing.

Five μg of a plasmid containing the polynucleotide is co-transfected with 1.0 μg of a commercially available linearized baculovirus DNA ("BaculoGold™ baculovirus DNA", Pharmingen, San Diego, CA), using the lipofection method described by Felgner et al., Proc. Natl. Acad. Sci. USA 84:7413-7417 (1987). One μg of BaculoGold™ virus DNA and 5 μg of the plasmid are mixed in a sterile well of a microtiter plate containing 50 μl of serum-free Grace's medium (Life Technologies Inc., Gaithersburg, MD). Afterwards, 10 μl Lipofectin plus 90 μl Grace's medium are added. mixed and incubated for 15 minutes at room temperature. Then the transfection mixture is added drop-wise to Sf9 insect cells (ATCC CRL 1711) seeded in a 35 mm tissue culture plate with 1 ml Grace's medium without serum. The plate is then incubated for 5 hours at 27° C. The transfection solution is then removed from the plate and 1 ml of Grace's insect medium supplemented with 10% fetal calf serum is added. Cultivation is then continued at 27° C for four days.

After four days the supernatant is collected and a plaque assay is performed, as described by Summers and Smith, *supra*. An agarose gel with "Blue Gal" (Life

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Technologies Inc., Gaithersburg) is used to allow easy identification and isolation of gal-expressing clones, which produce blue-stained plaques. (A detailed description of a "plaque assay" of this type can also be found in the user's guide for insect cell culture and baculovirology distributed by Life Technologies Inc., Gaithersburg, page 9-10.) After appropriate incubation, blue stained plaques are picked with the tip of a micropipettor (e.g., Eppendorf). The agar containing the recombinant viruses is then resuspended in a microcentrifuge tube containing 200 µl of Grace's medium and the suspension containing the recombinant baculovirus is used to infect Sf9 cells seeded in 35 mm dishes. Four days later the supernatants of these culture dishes are harvested and then they are stored at 4° C.

To verify the expression of the polypeptide, Sf9 cells are grown in Grace's medium supplemented with 10% heat-inactivated FBS. The cells are infected with the recombinant baculovirus containing the polynucleotide at a multiplicity of infection ("MOI") of about 2. If radiolabeled proteins are desired, 6 hours later the medium is removed and is replaced with SF900 II medium minus methionine and cysteine (available from Life Technologies Inc., Rockville, MD). After 42 hours, 5 μCi of ³⁵S-methionine and 5 μCi ³⁵S-cysteine (available from Amersham) are added. The cells are further incubated for 16 hours and then are harvested by centrifugation. The proteins in the supernatant as well as the intracellular proteins are analyzed by SDS-PAGE followed by autoradiography (if radiolabeled).

Microsequencing of the amino acid sequence of the amino terminus of purified protein may be used to determine the amino terminal sequence of the produced protein.

Example 8: Expression of a Polypeptide in Mammalian Cells

The polypeptide of the present invention can be expressed in a mammalian cell. A typical mammalian expression vector contains a promoter element, which mediates the initiation of transcription of mRNA, a protein coding sequence, and signals required for the termination of transcription and polyadenylation of the transcript. Additional elements include enhancers, Kozak sequences and intervening sequences flanked by donor and acceptor sites for RNA splicing. Highly efficient transcription is achieved with the early and late promoters from SV40, the long terminal repeats (LTRs) from Retroviruses, e.g., RSV, HTLVI, HIVI and the early promoter of the cytomegalovirus (CMV). However, cellular elements can also be used (e.g., the human actin promoter).

Suitable expression vectors for use in practicing the present invention include, for example, vectors such as pSVL and pMSG (Pharmacia, Uppsala, Sweden), pRSVcat (ATCC 37152), pSV2dhfr (ATCC 37146), pBC12MI (ATCC 67109), pCMVSport 2.0, and pCMVSport 3.0. Mammalian host cells that could be used

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include, human Hela, 293, H9 and Jurkat cells, mouse NIH3T3 and C127 cells, Cos 1, Cos 7 and CV1, quail QC1-3 cells, mouse L cells and Chinese hamster ovary (CHO) cells.

Alternatively, the polypeptide can be expressed in stable cell lines containing the polynucleotide integrated into a chromosome. The co-transfection with a selectable marker such as dhfr, gpt, neomycin, hygromycin allows the identification and isolation of the transfected cells.

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The transfected gene can also be amplified to express large amounts of the encoded protein. The DHFR (dihydrofolate reductase) marker is useful in developing cell lines that carry several hundred or even several thousand copies of the gene of interest. (See, e.g., Alt, F. W., et al., J. Biol. Chem. 253:1357-1370 (1978); Hamlin, J. L. and Ma, C., Biochem. et Biophys. Acta, 1097:107-143 (1990); Page, M. J. and Sydenham, M. A., Biotechnology 9:64-68 (1991).) Another useful selection marker is the enzyme glutamine synthase (GS) (Murphy et al., Biochem J. 227:277-279 (1991); Bebbington et al., Bio/Technology 10:169-175 (1992). Using these markers, the mammalian cells are grown in selective medium and the cells with the highest resistance are selected. These cell lines contain the amplified gene(s) integrated into a chromosome. Chinese hamster ovary (CHO) and NSO cells are often used for the production of proteins.

Derivatives of the plasmid pSV2-dhfr (ATCC Accession No. 37146), the expression vectors pC4 (ATCC Accession No. 209646) and pC6 (ATCC Accession No.209647) contain the strong promoter (LTR) of the Rous Sarcoma Virus (Cullen et al., Molecular and Cellular Biology, 438-447 (March, 1985)) plus a fragment of the CMV-enhancer (Boshart et al., Cell 41:521-530 (1985).) Multiple cloning sites, e.g., with the restriction enzyme cleavage sites BamHI, XbaI and Asp718, facilitate the cloning of the gene of interest. The vectors also contain the 3' intron, the polyadenylation and termination signal of the rat preproinsulin gene, and the mouse DHFR gene under control of the SV40 early promoter.

Specifically, the plasmid pC6, for example, is digested with appropriate restriction enzymes and then dephosphorylated using calf intestinal phosphates by procedures known in the art. The vector is then isolated from a 1% agarose gel.

A polynucleotide of the present invention is amplified according to the protocol outlined in Example 1. If the naturally occurring signal sequence is used to produce the secreted protein, the vector does not need a second signal peptide. Alternatively, if the naturally occurring signal sequence is not used, the vector can be modified to include a heterologous signal sequence. (See, e.g., WO 96/34891.)

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The amplified fragment is isolated from a 1% agarose gel using a commercially available kit ("Geneclean," BIO 101 Inc., La Jolla, Ca.). The fragment then is digested with appropriate restriction enzymes and again purified on a 1% agarose gel.

The amplified fragment is then digested with the same restriction enzyme and purified on a 1% agarose gel. The isolated fragment and the dephosphorylated vector are then ligated with T4 DNA ligase. *E. coli* HB101 or XL-1 Blue cells are then transformed and bacteria are identified that contain the fragment inserted into plasmid pC6 using, for instance, restriction enzyme analysis.

Chinese hamster ovary cells lacking an active DHFR gene is used for transfection. Five µg of the expression plasmid pC6 is cotransfected with 0.5 µg of the plasmid pSVneo using lipofectin (Felgner et al., supra). The plasmid pSV2-neo contains a dominant selectable marker, the neo gene from Tn5 encoding an enzyme that confers resistance to a group of antibiotics including G418. The cells are seeded in alpha minus MEM supplemented with 1 mg/ml G418. After 2 days, the cells are trypsinized and seeded in hybridoma cloning plates (Greiner, Germany) in alpha minus MEM supplemented with 10, 25, or 50 ng/ml of metothrexate plus 1 mg/ml G418. After about 10-14 days single clones are trypsinized and then seeded in 6-well petri dishes or 10 ml flasks using different concentrations of methotrexate (50 nM, 100 nM, 200 nM, 400 nM, 800 nM). Clones growing at the highest concentrations of methotrexate are then transferred to new 6-well plates containing even higher concentrations of methotrexate (1 μ M, 2 μ M, 5 μ M, 10 mM, 20 mM). The same procedure is repeated until clones are obtained which grow at a concentration of 100 -200 µM. Expression of the desired gene product is analyzed, for instance, by SDS-PAGE and Western blot or by reversed phase HPLC analysis.

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Example 9: Protein Fusions

The polypeptides of the present invention are preferably fused to other proteins. These fusion proteins can be used for a variety of applications. For example, fusion of the present polypeptides to His-tag, HA-tag, protein A, IgG domains, and maltose binding protein facilitates purification. (See Example 5; see also EP A 394,827; Traunecker, et al., Nature 331:84-86 (1988).) Similarly, fusion to IgG-1, IgG-3, and albumin increases the halflife time in vivo. Nuclear localization signals fused to the polypeptides of the present invention can target the protein to a specific subcellular localization, while covalent heterodimer or homodimers can increase or decrease the activity of a fusion protein. Fusion proteins can also create chimeric molecules having more than one function. Finally, fusion proteins can increase solubility and/or stability of the fused protein compared to the non-fused protein. All of the types of fusion

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proteins described above can be made by modifying the following protocol, which outlines the fusion of a polypeptide to an IgG molecule, or the protocol described in Example 5.

Briefly, the human Fc portion of the IgG molecule can be PCR amplified, using primers that span the 5' and 3' ends of the sequence described below. These primers also should have convenient restriction enzyme sites that will facilitate cloning into an expression vector, preferably a mammalian expression vector.

For example, if pC4 (Accession No. 209646) is used, the human Fc portion can be ligated into the BamHI cloning site. Note that the 3' BamHI site should be destroyed. Next, the vector containing the human Fc portion is re-restricted with BamHI, linearizing the vector, and a polynucleotide of the present invention, isolated by the PCR protocol described in Example 1, is ligated into this BamHI site. Note that the polynucleotide is cloned without a stop codon, otherwise a fusion protein will not be produced.

If the naturally occurring signal sequence is used to produce the secreted protein, pC4 does not need a second signal peptide. Alternatively, if the naturally occurring signal sequence is not used, the vector can be modified to include a heterologous signal sequence. (See, e.g., WO 96/34891.)

20 Human IgG Fc region:

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GGGATCCGGAGCCCAAATCTTCTGACAAAACTCACACATGCCCACCGTGCC
CAGCACCTGAATTCGAGGGTGCACCGTCAGTCTTCCTCTTCCCCCCAAAACC
CAAGGACACCCTCATGATCTCCCGGACTCCTGAGGTCACATGCGTGGTGT
GGACGTAAGCCACGAAGACCCTGAGGTCAAGTTCAACTGGTACGTGGACG
GCGTGGAGGTGCATAATGCCAAGACAAAGCCGCGGGAGGAGCAGTACAAC
AGCACGTACCGTGTGGTCAGCGTCCTCACCGTCCTGCACCAGGACTGGCTG
AATGGCAAGGAGTACAAGTGCAAGGTCTCCAACAAAGCCCTCCCAACCCCC
ATCGAGAAAACCATCTCCAAAGCCAAAGGGCAGCCCCGAGAACCACAGGT
GTACACCCTGCCCCCATCCCGGGATGAGCTGACCAAGAACCAGGTCAGCCT
GACCTGCCTGGTCAAAGGCTTCTATCCAAGCGACATCGCCGTGGAGTGGGA
GAGCAATGGGCAGCCGGAGAACAACTACAAGACCACGCCTCCCGTGCTGG
ACTCCGACGGCTCCTTCTTCCTCTACAGCAAGCTCACCGTGGACAAGAGCA
GGTGGCAGCAGGGGAACGTCTTCTCATGCTCCGTGATGCATGAGGCTCTGC
ACAACCACTACACGCAGAAGAGCCTCTCCCTGTCTCCGGGTAAATGAGTGC
GACGGCCGCGACTCTAGAGGAT (SEQ ID NO:1)

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Example 10: Production of an Antibody from a Polypeptide

The antibodies of the present invention can be prepared by a variety of methods. (See, Current Protocols, Chapter 2.) For example, cells expressing a polypeptide of the present invention is administered to an animal to induce the production of sera containing polyclonal antibodies. In a preferred method, a preparation of the secreted protein is prepared and purified to render it substantially free of natural contaminants. Such a preparation is then introduced into an animal in order to produce polyclonal antisera of greater specific activity.

In the most preferred method, the antibodies of the present invention are monoclonal antibodies (or protein binding fragments thereof). Such monoclonal antibodies can be prepared using hybridoma technology. (Köhler et al., Nature 256:495 (1975); Köhler et al., Eur. J. Immunol. 6:511 (1976); Köhler et al., Eur. J. Immunol. 6:292 (1976); Hammerling et al., in: Monoclonal Antibodies and T-Cell Hybridomas, Elsevier, N.Y., pp. 563-681 (1981).) In general, such procedures involve immunizing an animal (preferably a mouse) with polypeptide or, more preferably, with a secreted polypeptide-expressing cell. Such cells may be cultured in any suitable tissue culture medium; however, it is preferable to culture cells in Earle's modified Eagle's medium supplemented with 10% fetal bovine serum (inactivated at about 56°C), and supplemented with about 10 g/l of nonessential amino acids, about 1,000 U/ml of penicillin, and about 100 μg/ml of streptomycin.

The splenocytes of such mice are extracted and fused with a suitable myeloma cell line. Any suitable myeloma cell line may be employed in accordance with the present invention; however, it is preferable to employ the parent myeloma cell line (SP2O), available from the ATCC. After fusion, the resulting hybridoma cells are selectively maintained in HAT medium, and then cloned by limiting dilution as described by Wands et al. (Gastroenterology 80:225-232 (1981).) The hybridoma cells obtained through such a selection are then assayed to identify clones which secrete antibodies capable of binding the polypeptide.

Alternatively, additional antibodies capable of binding to the polypeptide can be produced in a two-step procedure using anti-idiotypic antibodies. Such a method makes use of the fact that antibodies are themselves antigens, and therefore, it is possible to obtain an antibody which binds to a second antibody. In accordance with this method, protein specific antibodies are used to immunize an animal, preferably a mouse. The splenocytes of such an animal are then used to produce hybridoma cells, and the hybridoma cells are screened to identify clones which produce an antibody whose ability to bind to the protein-specific antibody can be blocked by the polypeptide.

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Such antibodies comprise anti-idiotypic antibodies to the protein-specific antibody and can be used to immunize an animal to induce formation of further protein-specific antibodies.

It will be appreciated that Fab and F(ab')2 and other fragments of the antibodies of the present invention may be used according to the methods disclosed herein. Such fragments are typically produced by proteolytic cleavage, using enzymes such as papain (to produce Fab fragments) or pepsin (to produce F(ab')2 fragments). Alternatively, secreted protein-binding fragments can be produced through the application of recombinant DNA technology or through synthetic chemistry.

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For in vivo use of antibodies in humans, it may be preferable to use "humanized" chimeric monoclonal antibodies. Such antibodies can be produced using genetic constructs derived from hybridoma cells producing the monoclonal antibodies described above. Methods for producing chimeric antibodies are known in the art. (See, for review, Morrison, Science 229:1202 (1985); Oi et al., BioTechniques 4:214 (1986); Cabilly et al., U.S. Patent No. 4,816,567; Taniguchi et al., EP 171496; Morrison et al., EP 173494; Neuberger et al., WO 8601533; Robinson et al., WO 8702671; Boulianne et al., Nature 312:643 (1984); Neuberger et al., Nature 314:268 (1985).)

20 Example 11: Production Of Secreted Protein For High-Throughput Screening Assavs

The following protocol produces a supernatant containing a polypeptide to be tested. This supernatant can then be used in the Screening Assays described in Examples 13-20.

First, dilute Poly-D-Lysine (644 587 Boehringer-Mannheim) stock solution (1mg/ml in PBS) 1:20 in PBS (w/o calcium or magnesium 17-516F Biowhittaker) for a working solution of 50ug/ml. Add 200 ul of this solution to each well (24 well plates) and incubate at RT for 20 minutes. Be sure to distribute the solution over each well (note: a 12-channel pipetter may be used with tips on every other channel). Aspirate off the Poly-D-Lysine solution and rinse with 1ml PBS (Phosphate Buffered Saline). The PBS should remain in the well until just prior to plating the cells and plates may be poly-lysine coated in advance for up to two weeks.

Plate 293T cells (do not carry cells past P+20) at 2 x 10⁵ cells/well in .5ml DMEM(Dulbecco's Modified Eagle Medium)(with 4.5 G/L glucose and L-glutamine (12-604F Biowhittaker))/10% heat inactivated FBS(14-503F Biowhittaker)/1x Penstrep(17-602E Biowhittaker). Let the cells grow overnight.

The next day, mix together in a sterile solution basin: 300 ul Lipofectamine (18324-012 Gibco/BRL) and 5ml Optimem I (31985070 Gibco/BRL)/96-well plate. With a small volume multi-channel pipetter, aliquot approximately 2ug of an expression vector containing a polynucleotide insert, produced by the methods described in Examples 8 or 9, into an appropriately labeled 96-well round bottom plate. With a multi-channel pipetter, add 50ul of the Lipofectamine/Optimem I mixture to each well. Pipette up and down gently to mix. Incubate at RT 15-45 minutes. After about 20 minutes, use a multi-channel pipetter to add 150ul Optimem I to each well. As a control, one plate of vector DNA lacking an insert should be transfected with each set of transfections.

Preferably, the transfection should be performed by tag-teaming the following tasks. By tag-teaming, hands on time is cut in half, and the cells do not spend too much time on PBS. First, person A aspirates off the media from four 24-well plates of cells, and then person B rinses each well with .5-1ml PBS. Person A then aspirates off PBS rinse, and person B, using a12-channel pipetter with tips on every other channel, adds the 200ul of DNA/Lipofectamine/Optimem I complex to the odd wells first, then to the even wells, to each row on the 24-well plates. Incubate at 37°C for 6 hours.

While cells are incubating, prepare appropriate media, either 1%BSA in DMEM with 1x penstrep, or CHO-5 media (116.6 mg/L of CaCl2 (anhyd); 0.00130 mg/L 20 $CuSO_4-5H_2O$; 0.050 mg/L of Fe(NO₃)₃-9H₂O; 0.417 mg/L of FeSO₄-7H₂O; 311.80 mg/L of Kcl; 28.64 mg/L of MgCl₂; 48.84 mg/L of MgSO₄; 6995.50 mg/L of NaCl; 2400.0 mg/L of NaHCO₃; 62.50 mg/L of NaH₂PO₄-H₂0; 71.02 mg/L of Na₂HPO4; .4320 mg/L of ZnSO₄-7H₂O; .002 mg/L of Arachidonic Acid; 1.022 mg/L of Cholesterol; .070 mg/L of DL-alpha-Tocopherol-Acetate; 0.0520 mg/L of Linoleic 25 Acid; 0.010 mg/L of Linolenic Acid; 0.010 mg/L of Myristic Acid; 0.010 mg/L of Oleic Acid; 0.010 mg/L of Palmitric Acid; 0.010 mg/L of Palmitic Acid; 100 mg/L of Pluronic F-68; 0.010 mg/L of Stearic Acid; 2.20 mg/L of Tween 80; 4551 mg/L of D-Glucose; 130.85 mg/ml of L- Alanine; 147.50 mg/ml of L-Arginine-HCL; 7.50 mg/ml of L-Asparagine-H₂0; 6.65 mg/ml of L-Aspartic Acid; 29.56 mg/ml of L-Cystine-2HCL-H₂0; 31.29 mg/ml of L-Cystine-2HCL; 7.35 mg/ml of L-Glutamic Acid; 365.0 30 mg/ml of L-Glutamine; 18.75 mg/ml of Glycine; 52.48 mg/ml of L-Histidine-HCL-H₂0; 106.97 mg/ml of L-Isoleucine; 111.45 mg/ml of L-Leucine; 163.75 mg/ml of L-Lysine HCL; 32.34 mg/ml of L-Methionine; 68.48 mg/ml of L-Phenylalainine; 40.0 mg/ml of L-Proline; 26.25 mg/ml of L-Serine; 101.05 mg/ml of L-Threonine; 19.22 35 mg/ml of L-Tryptophan; 91.79 mg/ml of L-Tryrosine-2Na-2H₂0; 99.65 mg/ml of L-Valine; 0.0035 mg/L of Biotin; 3.24 mg/L of D-Ca Pantothenate; 11.78 mg/L of

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Choline Chloride; 4.65 mg/L of Folic Acid; 15.60 mg/L of i-Inositol; 3.02 mg/L of Niacinamide; 3.00 mg/L of Pyridoxal HCL; 0.031 mg/L of Pyridoxine HCL; 0.319 mg/L of Riboflavin; 3.17 mg/L of Thiamine HCL; 0.365 mg/L of Thymidine; and 0.680 mg/L of Vitamin B₁₂; 25 mM of HEPES Buffer; 2.39 mg/L of Na Hypoxanthine; 0.105 mg/L of Lipoic Acid; 0.081 mg/L of Sodium Putrescine-2HCL; 55.0 mg/L of Sodium Pyruvate; 0.0067 mg/L of Sodium Selenite; 20uM of Ethanolamine; 0.122 mg/L of Ferric Citrate; 41.70 mg/L of Methyl-B-Cyclodextrin complexed with Linoleic Acid; 33.33 mg/L of Methyl-B-Cyclodextrin complexed with Oleic Acid; and 10 mg/L of Methyl-B-Cyclodextrin complexed with Retinal) with 2mm glutamine and 1x penstrep. (BSA (81-068-3 Bayer) 100gm dissolved in 1L DMEM for a 10% BSA stock solution). Filter the media and collect 50 ul for endotoxin assay in 15ml polystyrene conical.

The transfection reaction is terminated, preferably by tag-teaming, at the end of the incubation period. Person A aspirates off the transfection media, while person B adds 1.5ml appropriate media to each well. Incubate at 37°C for 45 or 72 hours depending on the media used: 1%BSA for 45 hours or CHO-5 for 72 hours.

On day four, using a 300ul multichannel pipetter, aliquot 600ul in one 1ml deep well plate and the remaining supernatant into a 2ml deep well. The supernatants from each well can then be used in the assays described in Examples 13-20.

It is specifically understood that when activity is obtained in any of the assays described below using a supernatant, the activity originates from either the polypeptide directly (e.g., as a secreted protein) or by the polypeptide inducing expression of other proteins, which are then secreted into the supernatant. Thus, the invention further provides a method of identifying the protein in the supernatant characterized by an activity in a particular assay.

Example 12: Construction of GAS Reporter Construct

One signal transduction pathway involved in the differentiation and proliferation of cells is called the Jaks-STATs pathway. Activated proteins in the Jaks-STATs pathway bind to gamma activation site "GAS" elements or interferon-sensitive responsive element ("ISRE"), located in the promoter of many genes. The binding of a protein to these elements alter the expression of the associated gene.

GAS and ISRE elements are recognized by a class of transcription factors called Signal Transducers and Activators of Transcription, or "STATs." There are six members of the STATs family. Stat1 and Stat3 are present in many cell types, as is Stat2 (as response to IFN-alpha is widespread). Stat4 is more restricted and is not in

many cell types though it has been found in T helper class I, cells after treatment with IL-12. Stat5 was originally called mammary growth factor, but has been found at higher concentrations in other cells including myeloid cells. It can be activated in tissue culture cells by many cytokines.

The STATs are activated to translocate from the cytoplasm to the nucleus upon tyrosine phosphorylation by a set of kinases known as the Janus Kinase ("Jaks") family. Jaks represent a distinct family of soluble tyrosine kinases and include Tyk2, Jak1, Jak2, and Jak3. These kinases display significant sequence similarity and are generally catalytically inactive in resting cells.

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The Jaks are activated by a wide range of receptors summarized in the Table below. (Adapted from review by Schidler and Darnell, Ann. Rev. Biochem. 64:621-51 (1995).) A cytokine receptor family, capable of activating Jaks, is divided into two groups: (a) Class 1 includes receptors for IL-2, IL-3, IL-4, IL-6, IL-7, IL-9, IL-11, IL-12, IL-15, Epo, PRL, GH, G-CSF, GM-CSF, LIF, CNTF, and thrombopoietin: and (b) Class 2 includes IFN-a, IFN-g, and IL-10. The Class 1 receptors share a conserved cysteine motif (a set of four conserved cysteines and one tryptophan) and a WSXWS motif (a membrane proxial region encoding Trp-Ser-Xxx-Trp-Ser (SEQ ID NO:2)).

Thus, on binding of a ligand to a receptor, Jaks are activated, which in turn activate STATs, which then translocate and bind to GAS elements. This entire process is encompassed in the Jaks-STATs signal transduction pathway.

Therefore, activation of the Jaks-STATs pathway, reflected by the binding of the GAS or the ISRE element, can be used to indicate proteins involved in the proliferation and differentiation of cells. For example, growth factors and cytokines are known to activate the Jaks-STATs pathway. (See Table below.) Thus, by using GAS elements linked to reporter molecules, activators of the Jaks-STATs pathway can be identified.

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	Ligand	tyk2	JAKs Jakl	Jak2	Jak3	<u>STATS</u>	GAS(elements) or ISRE
5	IFN family IFN-a/B IFN-g Il-10	+ +.	+ + ?	- + ?	- -	1,2,3 1 1,3	ISRE GAS (IRF1>Lys6>IFP)
10	gp130 family IL-6 (Pleiotrohic) Il-11(Pleiotrohic) OnM(Pleiotrohic)	+ ? ? ?	+ + +	+ ? + +	????	1,3 1,3 1,3 1,3	GAS (IRF1>Lys6>IFP)
15	LIF(Pleiotrohic) CNTF(Pleiotrohic) G-CSF(Pleiotrohic) IL-12(Pleiotrohic)	: -/+ ? +	+++	+ + ? +	? ? +	1,3 1,3 1,3	
20	g-C family IL-2 (lymphocytes) IL-4 (lymph/myeloid) IL-7 (lymphocytes) IL-9 (lymphocytes) IL-13 (lymphocyte) IL-15	- - - - ?	+ + + + + + +	- - - - ?	+ + + + ? +	1,3,5 6 5 5 6 5	GAS GAS (IRF1 = IFP >>Ly6)(IgH) GAS GAS GAS GAS GAS
25	gp140 family IL-3 (myeloid) IL-5 (myeloid) GM-CSF (myeloid)	-	- - -	++++	- -	5 5 5	GAS (IRF1>IFP>>Ly6) GAS GAS
30	Growth hormone fam GH PRL EPO	ily ? ? ?	- +/- -	+ + +	- -	5 1,3,5 5	GAS(B-CAS>IRF1=IFP>>Ly6)
35 40	Receptor Tyrosine Ki EGF PDGF CSF-1	nases ? ? ?	+++++	+ + +	- - -	1,3 1,3 1,3	GAS (IRF1) GAS (not IRF1)

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To construct a synthetic GAS containing promoter element, which is used in the Biological Assays described in Examples 13-14, a PCR based strategy is employed to generate a GAS-SV40 promoter sequence. The 5' primer contains four tandem copies of the GAS binding site found in the IRF1 promoter and previously demonstrated to bind STATs upon induction with a range of cytokines (Rothman et al., Immunity 1:457-468 (1994).), although other GAS or ISRE elements can be used instead. The 5' primer also contains 18bp of sequence complementary to the SV40 early promoter sequence and is flanked with an XhoI site. The sequence of the 5' primer is: 5':GCGCCTCGAGATTTCCCCGAAATCTAGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATTAG:3' (SEQ ID NO:3)

The downstream primer is complementary to the SV40 promoter and is flanked with a Hind III site: 5':GCGGCAAGCTTTTTGCAAAGCCTAGGC:3' (SEQ ID NO:4)

PCR amplification is performed using the SV40 promoter template present in the B-gal:promoter plasmid obtained from Clontech. The resulting PCR fragment is digested with XhoI/Hind III and subcloned into BLSK2-. (Stratagene.) Sequencing with forward and reverse primers confirms that the insert contains the following sequence:

5':CTCGAGATTTCCCCGAAATCTAGATTTCCCCGAAATGATTTCCCCGAAATG
ATTTCCCCGAAATATCTGCCATCTCAATTAGTCAGCAACCATAGTCCCGCCC
CTAACTCCGCCCATCCCGCCCCTAACTCCGCCCAGTTCCGCCCATTCTCCGC
CCCATGGCTGACTAATTTTTTTTATTTATTTATGCAGAGGCCGAGGCCGCCTCGGC
CTCTGAGCTATTCCAGAAGTAGTGAGGAGGCTTTTTTGGAGGCCTAGGCTTT
TGCAAAAAGCTT:3' (SEQ ID NO:5)

With this GAS promoter element linked to the SV40 promoter, a GAS:SEAP2 reporter construct is next engineered. Here, the reporter molecule is a secreted alkaline phosphatase, or "SEAP." Clearly, however, any reporter molecule can be instead of SEAP, in this or in any of the other Examples. Well known reporter molecules that can be used instead of SEAP include chloramphenicol acetyltransferase (CAT), luciferase, alkaline phosphatase, B-galactosidase, green fluorescent protein (GFP), or any protein detectable by an antibody.

The above sequence confirmed synthetic GAS-SV40 promoter element is subcloned into the pSEAP-Promoter vector obtained from Clontech using HindIII and XhoI, effectively replacing the SV40 promoter with the amplified GAS:SV40 promoter element, to create the GAS-SEAP vector. However, this vector does not contain a neomycin resistance gene, and therefore, is not preferred for mammalian expression systems.

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Thus, in order to generate mammalian stable cell lines expressing the GAS-SEAP reporter, the GAS-SEAP cassette is removed from the GAS-SEAP vector using SalI and NotI, and inserted into a backbone vector containing the neomycin resistance gene, such as pGFP-1 (Clontech), using these restriction sites in the multiple cloning site, to create the GAS-SEAP/Neo vector. Once this vector is transfected into mammalian cells, this vector can then be used as a reporter molecule for GAS binding as described in Examples 13-14.

Other constructs can be made using the above description and replacing GAS with a different promoter sequence. For example, construction of reporter molecules containing NFK-B and EGR promoter sequences are described in Examples 15 and 16. However, many other promoters can be substituted using the protocols described in these Examples. For instance, SRE, IL-2, NFAT, or Osteocalcin promoters can be substituted, alone or in combination (e.g., GAS/NF-KB/EGR, GAS/NF-KB, Il-2/NFAT, or NF-KB/GAS). Similarly, other cell lines can be used to test reporter construct activity, such as HELA (epithelial), HUVEC (endothelial), Reh (B-cell), Saos-2 (osteoblast), HUVAC (aortic), or Cardiomyocyte.

Example 13: High-Throughput Screening Assay for T-cell Activity.

The following protocol is used to assess T-cell activity by identifying factors, such as growth factors and cytokines, that may proliferate or differentiate T-cells. T-cell activity is assessed using the GAS/SEAP/Neo construct produced in Example 12. Thus, factors that increase SEAP activity indicate the ability to activate the Jaks-STATS signal transduction pathway. The T-cell used in this assay is Jurkat T-cells (ATCC Accession No. TIB-152), although Molt-3 cells (ATCC Accession No. CRL-1552) and Molt-4 cells (ATCC Accession No. CRL-1582) cells can also be used.

Jurkat T-cells are lymphoblastic CD4+ Th1 helper cells. In order to generate stable cell lines, approximately 2 million Jurkat cells are transfected with the GAS-SEAP/neo vector using DMRIE-C (Life Technologies)(transfection procedure described below). The transfected cells are seeded to a density of approximately 20,000 cells per well and transfectants resistant to 1 mg/ml genticin selected. Resistant colonies are expanded and then tested for their response to increasing concentrations of interferon gamma. The dose response of a selected clone is demonstrated.

Specifically, the following protocol will yield sufficient cells for 75 wells containing 200 ul of cells. Thus, it is either scaled up, or performed in multiple to generate sufficient cells for multiple 96 well plates. Jurkat cells are maintained in RPMI + 10% serum with 1%Pen-Strep. Combine 2.5 mls of OPTI-MEM (Life Technologies)

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with 10 ug of plasmid DNA in a T25 flask. Add 2.5 ml OPTI-MEM containing 50 ul of DMRIE-C and incubate at room temperature for 15-45 mins.

During the incubation period, count cell concentration, spin down the required number of cells (10⁷ per transfection), and resuspend in OPTI-MEM to a final concentration of 10⁷ cells/ml. Then add 1ml of 1 x 10⁷ cells in OPTI-MEM to T25 flask and incubate at 37°C for 6 hrs. After the incubation, add 10 ml of RPMI + 15% serum.

The Jurkat:GAS-SEAP stable reporter lines are maintained in RPMI + 10% serum, 1 mg/ml Genticin, and 1% Pen-Strep. These cells are treated with supernatants containing a polypeptide as produced by the protocol described in Example 11.

On the day of treatment with the supernatant, the cells should be washed and resuspended in fresh RPMI + 10% serum to a density of 500,000 cells per ml. The exact number of cells required will depend on the number of supernatants being screened. For one 96 well plate, approximately 10 million cells (for 10 plates, 100 million cells) are required.

Transfer the cells to a triangular reservoir boat, in order to dispense the cells into a 96 well dish, using a 12 channel pipette. Using a 12 channel pipette, transfer 200 ul of cells into each well (therefore adding 100, 000 cells per well).

After all the plates have been seeded, 50 ul of the supernatants are transferred directly from the 96 well plate containing the supernatants into each well using a 12 channel pipette. In addition, a dose of exogenous interferon gamma (0.1, 1.0, 10 ng) is added to wells H9, H10, and H11 to serve as additional positive controls for the assay.

The 96 well dishes containing Jurkat cells treated with supernatants are placed in an incubator for 48 hrs (note: this time is variable between 48-72 hrs). 35 ul samples from each well are then transferred to an opaque 96 well plate using a 12 channel pipette. The opaque plates should be covered (using sellophene covers) and stored at -20°C until SEAP assays are performed according to Example 17. The plates containing the remaining treated cells are placed at 4°C and serve as a source of material for repeating the assay on a specific well if desired.

As a positive control, 100 Unit/ml interferon gamma can be used which is known to activate Jurkat T cells. Over 30 fold induction is typically observed in the positive control wells.

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Example 14: High-Throughput Screening Assay Identifying Myeloid Activity

The following protocol is used to assess myeloid activity by identifying factors, such as growth factors and cytokines, that may proliferate or differentiate myeloid cells. Myeloid cell activity is assessed using the GAS/SEAP/Neo construct produced in Example 12. Thus, factors that increase SEAP activity indicate the ability to activate the Jaks-STATS signal transduction pathway. The myeloid cell used in this assay is U937, a pre-monocyte cell line, although TF-1, HL60, or KG1 can be used.

To transiently transfect U937 cells with the GAS/SEAP/Neo construct produced in Example 12, a DEAE-Dextran method (Kharbanda et. al., 1994, Cell Growth & Differentiation, 5:259-265) is used. First, harvest 2x10e⁷ U937 cells and wash with PBS. The U937 cells are usually grown in RPMI 1640 medium containing 10% heat-inactivated fetal bovine serum (FBS) supplemented with 100 units/ml penicillin and 100 mg/ml streptomycin.

Next, suspend the cells in 1 ml of 20 mM Tris-HCl (pH 7.4) buffer containing 0.5 mg/ml DEAE-Dextran, 8 ug GAS-SEAP2 plasmid DNA, 140 mM NaCl, 5 mM KCl, 375 uM Na₂HPO₄.7H₂O, 1 mM MgCl₂, and 675 uM CaCl₂. Incubate at 37°C for 45 min.

Wash the cells with RPMI 1640 medium containing 10% FBS and then resuspend in 10 ml complete medium and incubate at 37°C for 36 hr.

The GAS-SEAP/U937 stable cells are obtained by growing the cells in 400 ug/ml G418. The G418-free medium is used for routine growth but every one to two months, the cells should be re-grown in 400 ug/ml G418 for couple of passages.

These cells are tested by harvesting $1x10^8$ cells (this is enough for ten 96-well plates assay) and wash with PBS. Suspend the cells in 200 ml above described growth medium, with a final density of $5x10^5$ cells/ml. Plate 200 ul cells per well in the 96-well plate (or $1x10^5$ cells/well).

Add 50 ul of the supernatant prepared by the protocol described in Example 11. Incubate at 37°C for 48 to 72 hr. As a positive control, 100 Unit/ml interferon gamma can be used which is known to activate U937 cells. Over 30 fold induction is typically observed in the positive control wells. SEAP assay the supernatant according to the protocol described in Example 17.

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Example 15: High-Throughput Screening Assay Identifying Neuronal Activity.

When cells undergo differentiation and proliferation, a group of genes are activated through many different signal transduction pathways. One of these genes, EGR1 (early growth response gene 1), is induced in various tissues and cell types upon activation. The promoter of EGR1 is responsible for such induction. Using the EGR1 promoter linked to reporter molecules, activation of cells can be assessed.

Particularly, the following protocol is used to assess neuronal activity in PC12 cell lines. PC12 cells (rat phenochromocytoma cells) are known to proliferate and/or differentiate by activation with a number of mitogens, such as TPA (tetradecanoyl phorbol acetate), NGF (nerve growth factor), and EGF (epidermal growth factor). The EGR1 gene expression is activated during this treatment. Thus, by stably transfecting PC12 cells with a construct containing an EGR promoter linked to SEAP reporter, activation of PC12 cells can be assessed.

The EGR/SEAP reporter construct can be assembled by the following protocol. The EGR-1 promoter sequence (-633 to +1)(Sakamoto K et al., Oncogene 6:867-871 (1991)) can be PCR amplified from human genomic DNA using the following primers:

- 5' GCGCTCGAGGGATGACAGCGATAGAACCCCGG -3' (SEQ ID NO:6)
- 5' GCGAAGCTTCGCGACTCCCGGATCCGCCTC-3' (SEQ ID NO:7)

Using the GAS:SEAP/Neo vector produced in Example 12, EGR1 amplified product can then be inserted into this vector. Linearize the GAS:SEAP/Neo vector using restriction enzymes XhoI/HindIII, removing the GAS/SV40 stuffer. Restrict the EGR1 amplified product with these same enzymes. Ligate the vector and the EGR1 promoter.

To prepare 96 well-plates for cell culture, two mls of a coating solution (1:30 dilution of collagen type I (Upstate Biotech Inc. Cat#08-115) in 30% ethanol (filter sterilized)) is added per one 10 cm plate or 50 ml per well of the 96-well plate, and allowed to air dry for 2 hr.

PC12 cells are routinely grown in RPMI-1640 medium (Bio Whittaker) containing 10% horse serum (JRH BIOSCIENCES, Cat. # 12449-78P), 5% heatinactivated fetal bovine serum (FBS) supplemented with 100 units/ml penicillin and 100 ug/ml streptomycin on a precoated 10 cm tissue culture dish. One to four split is done every three to four days. Cells are removed from the plates by scraping and resuspended with pipetting up and down for more than 15 times.

Transfect the EGR/SEAP/Neo construct into PC12 using the Lipofectamine protocol described in Example 11. EGR-SEAP/PC12 stable cells are obtained by growing the cells in 300 ug/ml G418. The G418-free medium is used for routine

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growth but every one to two months, the cells should be re-grown in 300 ug/ml G418 for couple of passages.

To assay for neuronal activity, a 10 cm plate with cells around 70 to 80% confluent is screened by removing the old medium. Wash the cells once with PBS (Phosphate buffered saline). Then starve the cells in low serum medium (RPMI-1640 containing 1% horse serum and 0.5% FBS with antibiotics) overnight.

The next morning, remove the medium and wash the cells with PBS. Scrape off the cells from the plate, suspend the cells well in 2 ml low serum medium. Count the cell number and add more low serum medium to reach final cell density as $5x10^5$ cells/ml.

Add 200 ul of the cell suspension to each well of 96-well plate (equivalent to $1x10^5$ cells/well). Add 50 ul supernatant produced by Example 11, 37°C for 48 to 72 hr. As a positive control, a growth factor known to activate PC12 cells through EGR can be used, such as 50 ng/ul of Neuronal Growth Factor (NGF). Over fifty-fold induction of SEAP is typically seen in the positive control wells. SEAP assay the supernatant according to Example 17.

Example 16: High-Throughput Screening Assay for T-cell Activity

NF-kB (Nuclear Factor kB) is a transcription factor activated by a wide variety of agents including the inflammatory cytokines IL-1 and TNF, CD30 and CD40, lymphotoxin-alpha and lymphotoxin-beta, by exposure to LPS or thrombin, and by expression of certain viral gene products. As a transcription factor, NF-kB regulates the expression of genes involved in immune cell activation, control of apoptosis (NF-kB appears to shield cells from apoptosis), B and T-cell development, anti-viral and antimicrobial responses, and multiple stress responses.

In non-stimulated conditions, NF- κB is retained in the cytoplasm with I-κB (Inhibitor κB). However, upon stimulation, I- κB is phosphorylated and degraded, causing NF- κB to shuttle to the nucleus, thereby activating transcription of target genes. Target genes activated by NF- κB include IL-2, IL-6, GM-CSF, ICAM-1 and class 1 MHC.

Due to its central role and ability to respond to a range of stimuli, reporter constructs utilizing the NF-kB promoter element are used to screen the supernatants produced in Example 11. Activators or inhibitors of NF-kB would be useful in treating

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diseases. For example, inhibitors of NF-kB could be used to treat those diseases related to the acute or chronic activation of NF-kB, such as rheumatoid arthritis.

The downstream primer is complementary to the 3' end of the SV40 promoter and is flanked with a Hind III site:

5':GCGGCAAGCTTTTTGCAAAGCCTAGGC:3' (SEQ ID NO:4)

PCR amplification is performed using the SV40 promoter template present in the pB-gal:promoter plasmid obtained from Clontech. The resulting PCR fragment is digested with XhoI and Hind III and subcloned into BLSK2-. (Stratagene) Sequencing with the T7 and T3 primers confirms the insert contains the following sequence:

5':CTCGAGGGGACTTTCCCGGGGACTTTCCGGGGACTTTCC ATCTGCCATCTCAATTAGTCAGCAACCATAGTCCCGCCCCTAACTCCGCCCA TCCCGCCCCTAACTCCGCCCAGTTCCGCCCATTCTCCGCCCCATGGCTGACT AATTTTTTTATTTATGCAGAGGCCGAGGCCGCCTCGGCCTCTGAGCTATTC CAGAAGTAGTGAGGAGGCTTTTTTGGAGGCCTAGGCTTTTGCAAAAAGCTT: 3' (SEQ ID NO:10)

Next, replace the SV40 minimal promoter element present in the pSEAP2-promoter plasmid (Clontech) with this NF-κB/SV40 fragment using XhoI and HindIII. However, this vector does not contain a neomycin resistance gene, and therefore, is not preferred for mammalian expression systems.

In order to generate stable mammalian cell lines, the NF-κB/SV40/SEAP

cassette is removed from the above NF-κB/SEAP vector using restriction enzymes SalI and NotI, and inserted into a vector containing neomycin resistance. Particularly, the NF-κB/SV40/SEAP cassette was inserted into pGFP-1 (Clontech), replacing the GFP gene, after restricting pGFP-1 with SalI and NotI.

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Once NF-kB/SV40/SEAP/Neo vector is created, stable Jurkat T-cells are created and maintained according to the protocol described in Example 13. Similarly, the method for assaying supernatants with these stable Jurkat T-cells is also described in Example 13. As a positive control, exogenous TNF alpha (0.1,1, 10 ng) is added to wells H9, H10, and H11, with a 5-10 fold activation typically observed.

Example 17: Assay for SEAP Activity

As a reporter molecule for the assays described in Examples 13-16, SEAP activity is assayed using the Tropix Phospho-light Kit (Cat. BP-400) according to the following general procedure. The Tropix Phospho-light Kit supplies the Dilution, Assay, and Reaction Buffers used below.

Prime a dispenser with the 2.5x Dilution Buffer and dispense 15 µl of 2.5x dilution buffer into Optiplates containing 35 µl of a supernatant. Seal the plates with a plastic sealer and incubate at 65°C for 30 min. Separate the Optiplates to avoid uneven heating.

Cool the samples to room temperature for 15 minutes. Empty the dispenser and prime with the Assay Buffer. Add 50 µl Assay Buffer and incubate at room temperature 5 min. Empty the dispenser and prime with the Reaction Buffer (see the table below). Add 50 µl Reaction Buffer and incubate at room temperature for 20 minutes. Since the intensity of the chemiluminescent signal is time dependent, and it takes about 10 minutes to read 5 plates on luminometer, one should treat 5 plates at each time and start the second set 10 minutes later.

Read the relative light unit in the luminometer. Set H12 as blank, and print the results. An increase in chemiluminescence indicates reporter activity.

Reaction Buffer Formulation:

Reaction	Duiter Formulation:	
# of plates	Rxn buffer diluent (ml)	CSPD (ml)
10	60	3
11	65	3.25
12	70	3.5
13	75	3.75
14	80	4
15	85	4.25
16	90	4.5
17	95	4.75
18	100	5
19	105	5.25
20	110	5.5
21	115	5.75
22	120	. 6

23	125	6.25
24	130	6.5
25	135	6.75
26	140	7
27	145	7.25
28	150	7.5 .
29	155	7.75
30	160	8
31	165	8.25
32	170	8.5
33	175	8.75
34	180	9
35	185	9.25
36	190	9.5
37	195	9.75
38	200	10
39	205	10.25
40	210	10.5
41	215	10.75
42	220	11
43	225	11.25
44	230	11.5
45	235	11.75
46	240	12
47	245	12.25
48	250	12.5
49	255	12.75
50	260	13

Example 18: High-Throughput Screening Assay Identifying Changes in Small Molecule Concentration and Membrane Permeability

Binding of a ligand to a receptor is known to alter intracellular levels of small molecules, such as calcium, potassium, sodium, and pH, as well as alter membrane potential. These alterations can be measured in an assay to identify supernatants which bind to receptors of a particular cell. Although the following protocol describes an assay for calcium, this protocol can easily be modified to detect changes in potassium, sodium, pH, membrane potential, or any other small molecule which is detectable by a fluorescent probe.

The following assay uses Fluorometric Imaging Plate Reader ("FLIPR") to measure changes in fluorescent molecules (Molecular Probes) that bind small molecules. Clearly, any fluorescent molecule detecting a small molecule can be used instead of the calcium fluorescent molecule, fluo-3, used here.

For adherent cells, seed the cells at 10,000 -20,000 cells/well in a Co-star black 96-well plate with clear bottom. The plate is incubated in a CO_2 incubator for 20 hours. The adherent cells are washed two times in Biotek washer with 200 ul of HBSS (Hank's Balanced Salt Solution) leaving 100 ul of buffer after the final wash.

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A stock solution of 1 mg/ml fluo-3 is made in 10% pluronic acid DMSO. To load the cells with fluo-3, 50 ul of 12 ug/ml fluo-3 is added to each well. The plate is incubated at 37°C in a CO₂ incubator for 60 min. The plate is washed four times in the Biotek washer with HBSS leaving 100 ul of buffer.

For non-adherent cells, the cells are spun down from culture media. Cells are re-suspended to 2-5x10⁶ cells/ml with HBSS in a 50-ml conical tube. 4 ul of 1 mg/ml fluo-3 solution in 10% pluronic acid DMSO is added to each ml of cell suspension. The tube is then placed in a 37°C water bath for 30-60 min. The cells are washed twice with HBSS, resuspended to 1x10⁶ cells/ml, and dispensed into a microplate, 100 ul/well. The plate is centrifuged at 1000 rpm for 5 min. The plate is then washed once in Denley CellWash with 200 ul, followed by an aspiration step to 100 ul final volume.

For a non-cell based assay, each well contains a fluorescent molecule, such as fluo-3. The supernatant is added to the well, and a change in fluorescence is detected.

To measure the fluorescence of intracellular calcium, the FLIPR is set for the following parameters: (1) System gain is 300-800 mW; (2) Exposure time is 0.4 second; (3) Camera F/stop is F/2; (4) Excitation is 488 nm; (5) Emission is 530 nm; and (6) Sample addition is 50 ul. Increased emission at 530 nm indicates an extracellular signaling event which has resulted in an increase in the intracellular Ca⁺⁺ concentration.

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Example 19: High-Throughput Screening Assay Identifying Tyrosine Kinase Activity

The Protein Tyrosine Kinases (PTK) represent a diverse group of transmembrane and cytoplasmic kinases. Within the Receptor Protein Tyrosine Kinase RPTK) group are receptors for a range of mitogenic and metabolic growth factors including the PDGF, FGF, EGF, NGF, HGF and Insulin receptor subfamilies. In addition there are a large family of RPTKs for which the corresponding ligand is unknown. Ligands for RPTKs include mainly secreted small proteins, but also membrane-bound and extracellular matrix proteins.

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Activation of RPTK by ligands involves ligand-mediated receptor dimerization, resulting in transphosphorylation of the receptor subunits and activation of the cytoplasmic tyrosine kinases. The cytoplasmic tyrosine kinases include receptor associated tyrosine kinases of the src-family (e.g., src, yes, lck, lyn, fyn) and non-receptor linked and cytosolic protein tyrosine kinases, such as the Jak family, members of which mediate signal transduction triggered by the cytokine superfamily of receptors (e.g., the Interleukins, Interferons, GM-CSF, and Leptin).

Because of the wide range of known factors capable of stimulating tyrosine kinase activity, the identification of novel human secreted proteins capable of activating tyrosine kinase signal transduction pathways are of interest. Therefore, the following protocol is designed to identify those novel human secreted proteins capable of activating the tyrosine kinase signal transduction pathways.

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Seed target cells (e.g., primary keratinocytes) at a density of approximately 25,000 cells per well in a 96 well Loprodyne Silent Screen Plates purchased from Nalge Nunc (Naperville, IL). The plates are sterilized with two 30 minute rinses with 100% ethanol, rinsed with water and dried overnight. Some plates are coated for 2 hr with 100 ml of cell culture grade type I collagen (50 mg/ml), gelatin (2%) or polylysine (50 mg/ml), all of which can be purchased from Sigma Chemicals (St. Louis, MO) or 10% Matrigel purchased from Becton Dickinson (Bedford,MA), or calf serum, rinsed with PBS and stored at 4°C. Cell growth on these plates is assayed by seeding 5,000 cells/well in growth medium and indirect quantitation of cell number through use of alamarBlue as described by the manufacturer Alamar Biosciences, Inc. (Sacramento, CA) after 48 hr. Falcon plate covers #3071 from Becton Dickinson (Bedford,MA) are used to cover the Loprodyne Silent Screen Plates. Falcon Microtest III cell culture plates can also be used in some proliferation experiments.

To prepare extracts, A431 cells are seeded onto the nylon membranes of Loprodyne plates (20,000/200ml/well) and cultured overnight in complete medium. Cells are quiesced by incubation in serum-free basal medium for 24 hr. After 5-20 minutes treatment with EGF (60ng/ml) or 50 ul of the supernatant produced in Example 11, the medium was removed and 100 ml of extraction buffer ((20 mM HEPES pH 7.5, 0.15 M NaCl, 1% Triton X-100, 0.1% SDS, 2 mM Na3VO4, 2 mM Na4P2O7 and a cocktail of protease inhibitors (# 1836170) obtained from Boeheringer Mannheim (Indianapolis, IN) is added to each well and the plate is shaken on a rotating shaker for 5 minutes at 4°C. The plate is then placed in a vacuum transfer manifold and the extract filtered through the 0.45 mm membrane bottoms of each well using house vacuum. Extracts are collected in a 96-well catch/assay plate in the bottom of the vacuum manifold and immediately placed on ice. To obtain extracts clarified by centrifugation, the content of each well, after detergent solubilization for 5 minutes, is removed and centrifuged for 15 minutes at 4°C at 16,000 x g.

Test the filtered extracts for levels of tyrosine kinase activity. Although many methods of detecting tyrosine kinase activity are known, one method is described here.

Generally, the tyrosine kinase activity of a supernatant is evaluated by determining its ability to phosphorylate a tyrosine residue on a specific substrate (a

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biotinylated peptide). Biotinylated peptides that can be used for this purpose include PSK1 (corresponding to amino acids 6-20 of the cell division kinase cdc2-p34) and PSK2 (corresponding to amino acids 1-17 of gastrin). Both peptides are substrates for a range of tyrosine kinases and are available from Boehringer Mannheim.

The tyrosine kinase reaction is set up by adding the following components in order. First, add 10ul of 5uM Biotinylated Peptide, then 10ul ATP/Mg₂₊ (5mM ATP/50mM MgCl₂), then 10ul of 5x Assay Buffer (40mM imidazole hydrochloride, pH7.3, 40 mM beta-glycerophosphate, 1mM EGTA, 100mM MgCl₂, 5 mM MnCl₂, 0.5 mg/ml BSA), then 5ul of Sodium Vanadate(1mM), and then 5ul of water. Mix the components gently and preincubate the reaction mix at 30°C for 2 min. Initial the reaction by adding 10ul of the control enzyme or the filtered supernatant.

The tyrosine kinase assay reaction is then terminated by adding 10 ul of 120mm EDTA and place the reactions on ice.

Tyrosine kinase activity is determined by transferring 50 ul aliquot of reaction mixture to a microtiter plate (MTP) module and incubating at 37°C for 20 min. This allows the streptavadin coated 96 well plate to associate with the biotinylated peptide. Wash the MTP module with 300ul/well of PBS four times. Next add 75 ul of anti-phospotyrosine antibody conjugated to horse radish peroxidase(anti-P-Tyr-POD(0.5u/ml)) to each well and incubate at 37°C for one hour. Wash the well as above.

Next add 100ul of peroxidase substrate solution (Boehringer Mannheim) and incubate at room temperature for at least 5 mins (up to 30 min). Measure the absorbance of the sample at 405 nm by using ELISA reader. The level of bound peroxidase activity is quantitated using an ELISA reader and reflects the level of tyrosine kinase activity.

Example 20: High-Throughput Screening Assay Identifying Phosphorylation Activity

As a potential alternative and/or compliment to the assay of protein tyrosine kinase activity described in Example 19, an assay which detects activation (phosphorylation) of major intracellular signal transduction intermediates can also be used. For example, as described below one particular assay can detect tyrosine phosphorylation of the Erk-1 and Erk-2 kinases. However, phosphorylation of other molecules, such as Raf, JNK, p38 MAP, Map kinase kinase (MEK), MEK kinase, Src, Muscle specific kinase (MuSK), IRAK, Tec, and Janus, as well as any other

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phosphoserine, phosphotyrosine, or phosphothreonine molecule, can be detected by substituting these molecules for Erk-1 or Erk-2 in the following assay.

Specifically, assay plates are made by coating the wells of a 96-well ELISA plate with 0.1ml of protein G (1ug/ml) for 2 hr at room temp, (RT). The plates are then rinsed with PBS and blocked with 3% BSA/PBS for 1 hr at RT. The protein G plates are then treated with 2 commercial monoclonal antibodies (100ng/well) against Erk-1 and Erk-2 (1 hr at RT) (Santa Cruz Biotechnology). (To detect other molecules, this step can easily be modified by substituting a monoclonal antibody detecting any of the above described molecules.) After 3-5 rinses with PBS, the plates are stored at 4°C until use.

A431 cells are seeded at 20,000/well in a 96-well Loprodyne filterplate and cultured overnight in growth medium. The cells are then starved for 48 hr in basal medium (DMEM) and then treated with EGF (6ng/well) or 50 ul of the supernatants obtained in Example 11 for 5-20 minutes. The cells are then solubilized and extracts filtered directly into the assay plate.

After incubation with the extract for 1 hr at RT, the wells are again rinsed. As a positive control, a commercial preparation of MAP kinase (10ng/well) is used in place of A431 extract. Plates are then treated with a commercial polyclonal (rabbit) antibody (lug/ml) which specifically recognizes the phosphorylated epitope of the Erk-1 and Erk-2 kinases (1 hr at RT). This antibody is biotinylated by standard procedures. The bound polyclonal antibody is then quantitated by successive incubations with Europium-streptavidin and Europium fluorescence enhancing reagent in the Wallac DELFIA instrument (time-resolved fluorescence). An increased fluorescent signal over background indicates a phosphorylation.

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Example 21: Method of Determining Alterations in a Gene Corresponding to a Polynucleotide

RNA isolated from entire families or individual patients presenting with a phenotype of interest (such as a disease) is be isolated. cDNA is then generated from these RNA samples using protocols known in the art. (See, Sambrook.) The cDNA is then used as a template for PCR, employing primers surrounding regions of interest in SEQ ID NO:X. Suggested PCR conditions consist of 35 cycles at 95°C for 30 seconds; 60-120 seconds at 52-58°C; and 60-120 seconds at 70°C, using buffer solutions described in Sidransky, D., et al., Science 252:706 (1991).

PCR products are then sequenced using primers labeled at their 5' end with T4 polynucleotide kinase, employing SequiTherm Polymerase. (Epicentre Technologies).

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The intron-exon borders of selected exons is also determined and genomic PCR products analyzed to confirm the results. PCR products harboring suspected mutations is then cloned and sequenced to validate the results of the direct sequencing.

PCR products is cloned into T-tailed vectors as described in Holton, T.A. and Graham, M.W., Nucleic Acids Research, 19:1156 (1991) and sequenced with T7 polymerase (United States Biochemical). Affected individuals are identified by mutations not present in unaffected individuals.

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Genomic rearrangements are also observed as a method of determining alterations in a gene corresponding to a polynucleotide. Genomic clones isolated according to Example 2 are nick-translated with digoxigenindeoxy-uridine 5'-triphosphate (Boehringer Manheim), and FISH performed as described in Johnson, Cg. et al., Methods Cell Biol. 35:73-99 (1991). Hybridization with the labeled probe is carried out using a vast excess of human cot-1 DNA for specific hybridization to the corresponding genomic locus.

Chromosomes are counterstained with 4,6-diamino-2-phenylidole and propidium iodide, producing a combination of C- and R-bands. Aligned images for precise mapping are obtained using a triple-band filter set (Chroma Technology, Brattleboro, VT) in combination with a cooled charge-coupled device camera (Photometrics, Tucson, AZ) and variable excitation wavelength filters. (Johnson, Cv. et al., Genet. Anal. Tech. Appl., 8:75 (1991).) Image collection, analysis and chromosomal fractional length measurements are performed using the ISee Graphical Program System. (Inovision Corporation, Durham, NC.) Chromosome alterations of the genomic region hybridized by the probe are identified as insertions, deletions, and translocations. These alterations are used as a diagnostic marker for an associated disease.

Example 22: Method of Detecting Abnormal Levels of a Polypeptide in a Biological Sample

A polypeptide of the present invention can be detected in a biological sample, and if an increased or decreased level of the polypeptide is detected, this polypeptide is a marker for a particular phenotype. Methods of detection are numerous, and thus, it is understood that one skilled in the art can modify the following assay to fit their particular needs.

For example, antibody-sandwich ELISAs are used to detect polypeptides in a sample, preferably a biological sample. Wells of a microtiter plate are coated with specific antibodies, at a final concentration of 0.2 to 10 ug/ml. The antibodies are either monoclonal or polyclonal and are produced by the method described in Example 10.

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The wells are blocked so that non-specific binding of the polypeptide to the well is reduced.

The coated wells are then incubated for > 2 hours at RT with a sample containing the polypeptide. Preferably, serial dilutions of the sample should be used to validate results. The plates are then washed three times with deionized or distilled water to remove unbounded polypeptide.

Next, 50 ul of specific antibody-alkaline phosphatase conjugate, at a concentration of 25-400 ng, is added and incubated for 2 hours at room temperature. The plates are again washed three times with deionized or distilled water to remove unbounded conjugate.

Add 75 ul of 4-methylumbelliferyl phosphate (MUP) or p-nitrophenyl phosphate (NPP) substrate solution to each well and incubate 1 hour at room temperature. Measure the reaction by a microtiter plate reader. Prepare a standard curve, using serial dilutions of a control sample, and plot polypeptide concentration on the X-axis (log scale) and fluorescence or absorbance of the Y-axis (linear scale). Interpolate the concentration of the polypeptide in the sample using the standard curve.

Example 23: Formulating a Polypeptide

The secreted polypeptide composition will be formulated and dosed in a fashion consistent with good medical practice, taking into account the clinical condition of the individual patient (especially the side effects of treatment with the secreted polypeptide alone), the site of delivery, the method of administration, the scheduling of administration, and other factors known to practitioners. The "effective amount" for purposes herein is thus determined by such considerations.

As a general proposition, the total pharmaceutically effective amount of secreted polypeptide administered parenterally per dose will be in the range of about 1 μ g/kg/day to 10 mg/kg/day of patient body weight, although, as noted above, this will be subject to therapeutic discretion. More preferably, this dose is at least 0.01 mg/kg/day, and most preferably for humans between about 0.01 and 1 mg/kg/day for the hormone. If given continuously, the secreted polypeptide is typically administered at a dose rate of about 1 μ g/kg/hour to about 50 μ g/kg/hour, either by 1-4 injections per day or by continuous subcutaneous infusions, for example, using a mini-pump. An intravenous bag solution may also be employed. The length of treatment needed to observe changes and the interval following treatment for responses to occur appears to vary depending on the desired effect.

Pharmaceutical compositions containing the secreted protein of the invention are administered orally, rectally, parenterally, intracistemally, intravaginally,

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intraperitoneally, topically (as by powders, ointments, gels, drops or transdermal patch), bucally, or as an oral or nasal spray. "Pharmaceutically acceptable carrier" refers to a non-toxic solid, semisolid or liquid filler, diluent, encapsulating material or formulation auxiliary of any type. The term "parenteral" as used herein refers to modes of administration which include intravenous, intramuscular, intraperitoneal, intrasternal, subcutaneous and intraarticular injection and infusion.

The secreted polypeptide is also suitably administered by sustained-release systems. Suitable examples of sustained-release compositions include semi-permeable polymer matrices in the form of shaped articles, e.g., films, or mirocapsules. Sustained-release matrices include polylactides (U.S. Pat. No. 3,773,919, EP 58,481), 10 copolymers of L-glutamic acid and gamma-ethyl-L-glutamate (Sidman, U. et al., Biopolymers 22:547-556 (1983)), poly (2-hydroxyethyl methacrylate) (R. Langer et al., J. Biomed. Mater. Res. 15:167-277 (1981), and R. Langer, Chem. Tech. 12:98-22 105 (1982)), ethylene vinyl acetate (R. Langer et al.) or poly-D- (-)-3-hydroxybutyric acid (EP 133,988). Sustained-release compositions also include liposomally entrapped 15 polypeptides. Liposomes containing the secreted polypeptide are prepared by methods known per se: DE 3,218,121; Epstein et al., Proc. Natl. Acad. Sci. USA 82:3688-3692 (1985); Hwang et al., Proc. Natl. Acad. Sci. USA 77:4030-4034 (1980); EP 52,322; EP 36.676; EP 88,046; EP 143,949; EP 142,641; Japanese Pat. Appl. 83-118008; U.S. Pat. Nos. 4,485,045 and 4,544,545; and EP 102,324. Ordinarily, the liposomes 20 are of the small (about 200-800 Angstroms) unilamellar type in which the lipid content is greater than about 30 mol. percent cholesterol, the selected proportion being adjusted for the optimal secreted polypeptide therapy.

For parenteral administration, in one embodiment, the secreted polypeptide is formulated generally by mixing it at the desired degree of purity, in a unit dosage injectable form (solution, suspension, or emulsion), with a pharmaceutically acceptable carrier, i.e., one that is non-toxic to recipients at the dosages and concentrations employed and is compatible with other ingredients of the formulation. For example, the formulation preferably does not include oxidizing agents and other compounds that are known to be deleterious to polypeptides.

Generally, the formulations are prepared by contacting the polypeptide uniformly and intimately with liquid carriers or finely divided solid carriers or both. Then, if necessary, the product is shaped into the desired formulation. Preferably the carrier is a parenteral carrier, more preferably a solution that is isotonic with the blood of the recipient. Examples of such carrier vehicles include water, saline, Ringer's solution, and dextrose solution. Non-aqueous vehicles such as fixed oils and ethyl oleate are also useful herein, as well as liposomes.

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The carrier suitably contains minor amounts of additives such as substances that enhance isotonicity and chemical stability. Such materials are non-toxic to recipients at the dosages and concentrations employed, and include buffers such as phosphate, citrate, succinate, acetic acid, and other organic acids or their salts; antioxidants such as ascorbic acid; low molecular weight (less than about ten residues) polypeptides, e.g., polyarginine or tripeptides; proteins, such as serum albumin, gelatin, or immunoglobulins; hydrophilic polymers such as polyvinylpyrrolidone; amino acids, such as glycine, glutamic acid, aspartic acid, or arginine; monosaccharides, disaccharides, and other carbohydrates including cellulose or its derivatives, glucose, manose, or dextrins; chelating agents such as EDTA; sugar alcohols such as mannitol or sorbitol; counterions such as sodium; and/or nonionic surfactants such as polysorbates, poloxamers, or PEG.

The secreted polypeptide is typically formulated in such vehicles at a concentration of about 0.1 mg/ml to 100 mg/ml, preferably 1-10 mg/ml, at a pH of about 3 to 8. It will be understood that the use of certain of the foregoing excipients, carriers, or stabilizers will result in the formation of polypeptide salts.

Any polypeptide to be used for therapeutic administration can be sterile. Sterility is readily accomplished by filtration through sterile filtration membranes (e.g., 0.2 micron membranes). Therapeutic polypeptide compositions generally are placed into a container having a sterile access port, for example, an intravenous solution bag or vial having a stopper pierceable by a hypodermic injection needle.

Polypeptides ordinarily will be stored in unit or multi-dose containers, for example, sealed ampoules or vials, as an aqueous solution or as a lyophilized formulation for reconstitution. As an example of a lyophilized formulation, 10-ml vials are filled with 5 ml of sterile-filtered 1% (w/v) aqueous polypeptide solution, and the resulting mixture is lyophilized. The infusion solution is prepared by reconstituting the lyophilized polypeptide using bacteriostatic Water-for-Injection.

The invention also provides a pharmaceutical pack or kit comprising one or more containers filled with one or more of the ingredients of the pharmaceutical compositions of the invention. Associated with such container(s) can be a notice in the form prescribed by a governmental agency regulating the manufacture, use or sale of pharmaceuticals or biological products, which notice reflects approval by the agency of manufacture, use or sale for human administration. In addition, the polypeptides of the present invention may be employed in conjunction with other therapeutic compounds.

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Example 24: Method of Treating Decreased Levels of the Polypeptide

It will be appreciated that conditions caused by a decrease in the standard or normal expression level of a secreted protein in an individual can be treated by administering the polypeptide of the present invention, preferably in the secreted form. Thus, the invention also provides a method of treatment of an individual in need of an increased level of the polypeptide comprising administering to such an individual a pharmaceutical composition comprising an amount of the polypeptide to increase the activity level of the polypeptide in such an individual.

For example, a patient with decreased levels of a polypeptide receives a daily dose 0.1-100 ug/kg of the polypeptide for six consecutive days. Preferably, the polypeptide is in the secreted form. The exact details of the dosing scheme, based on administration and formulation, are provided in Example 23.

Example 25: Method of Treating Increased Levels of the Polypeptide

Antisense technology is used to inhibit production of a polypeptide of the present invention. This technology is one example of a method of decreasing levels of a polypeptide, preferably a secreted form, due to a variety of etiologies, such as cancer.

For example, a patient diagnosed with abnormally increased levels of a polypeptide is administered intravenously antisense polynucleotides at 0.5, 1.0, 1.5, 2.0 and 3.0 mg/kg day for 21 days. This treatment is repeated after a 7-day rest period if the treatment was well tolerated. The formulation of the antisense polynucleotide is provided in Example 23.

Example 26: Method of Treatment Using Gene Therapy

One method of gene therapy transplants fibroblasts, which are capable of expressing a polypeptide, onto a patient. Generally, fibroblasts are obtained from a subject by skin biopsy. The resulting tissue is placed in tissue-culture medium and separated into small pieces. Small chunks of the tissue are placed on a wet surface of a tissue culture flask, approximately ten pieces are placed in each flask. The flask is turned upside down, closed tight and left at room temperature over night. After 24 hours at room temperature, the flask is inverted and the chunks of tissue remain fixed to the bottom of the flask and fresh media (e.g., Ham's F12 media, with 10% FBS, penicillin and streptomycin) is added. The flasks are then incubated at 37°C for approximately one week.

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At this time, fresh media is added and subsequently changed every several days. After an additional two weeks in culture, a monolayer of fibroblasts emerge. The monolayer is trypsinized and scaled into larger flasks.

pMV-7 (Kirschmeier, P.T. et al., DNA, 7:219-25 (1988)), flanked by the long terminal repeats of the Moloney murine sarcoma virus, is digested with EcoRI and HindIII and subsequently treated with calf intestinal phosphatase. The linear vector is fractionated on agarose gel and purified, using glass beads.

The cDNA encoding a polypeptide of the present invention can be amplified using PCR primers which correspond to the 5' and 3' end sequences respectively as set forth in Example 1. Preferably, the 5' primer contains an EcoRI site and the 3' primer includes a HindIII site. Equal quantities of the Moloney murine sarcoma virus linear backbone and the amplified EcoRI and HindIII fragment are added together, in the presence of T4 DNA ligase. The resulting mixture is maintained under conditions appropriate for ligation of the two fragments. The ligation mixture is then used to transform bacteria HB101, which are then plated onto agar containing kanamycin for the purpose of confirming that the vector has the gene of interest properly inserted.

The amphotropic pA317 or GP+am12 packaging cells are grown in tissue culture to confluent density in Dulbecco's Modified Eagles Medium (DMEM) with 10% calf serum (CS), penicillin and streptomycin. The MSV vector containing the gene is then added to the media and the packaging cells transduced with the vector. The packaging cells now produce infectious viral particles containing the gene (the packaging cells are now referred to as producer cells).

Fresh media is added to the transduced producer cells, and subsequently, the media is harvested from a 10 cm plate of confluent producer cells. The spent media, containing the infectious viral particles, is filtered through a millipore filter to remove detached producer cells and this media is then used to infect fibroblast cells. Media is removed from a sub-confluent plate of fibroblasts and quickly replaced with the media from the producer cells. This media is removed and replaced with fresh media. If the titer of virus is high, then virtually all fibroblasts will be infected and no selection is required. If the titer is very low, then it is necessary to use a retroviral vector that has a selectable marker, such as neo or his. Once the fibroblasts have been efficiently infected, the fibroblasts are analyzed to determine whether protein is produced.

The engineered fibroblasts are then transplanted onto the host, either alone or after having been grown to confluence on cytodex 3 microcarrier beads.

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It will be clear that the invention may be practiced otherwise than as particularly described in the foregoing description and examples. Numerous modifications and variations of the present invention are possible in light of the above teachings and, therefore, are within the scope of the appended claims.

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The entire disclosure of each document cited (including patents, patent applications, journal articles, abstracts, laboratory manuals, books, or other disclosures) in the Background of the Invention, Detailed Description, and Examples is hereby incorporated herein by reference.

(1) GENERAL INFORMATION:

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(i) APPLICANT:
                           Human Genome Sciences, Inc. et al.
           (ii) TITLE OF INVENTION: 87 Human Secreted Proteins
5
           (iii) NUMBER OF SEQUENCES: 323
           (iv) CORRESPONDENCE ADDRESS:
                 (A) ADDRESSEE: Human Genome Sciences, Inc.
                 (B) STREET: 9410 Key West Avenue
10
                 (C) CITY: Rockville
                 (D) STATE: Maryland
                 (E) COUNTRY: USA
                 (F) ZIP: 20850
15
           (v) COMPUTER READABLE FORM:
                 (A) MEDIUM TYPE: Diskette, 3.50 inch, 1.4Mb storage
                 (B) COMPUTER: HP Vectra 486/33
20
                 (C) OPERATING SYSTEM: MSDOS version 6.2
                 (D) SOFTWARE: ASCII Text
           (vi) CURRENT APPLICATION DATA:
25
                  (A) APPLICATION NUMBER:
                  (B) FILING DATE: March 19, 1998
                  (C) CLASSIFICATION:
30
           (vii) PRIOR APPLICATION DATA:
                  (A) APPLICATION NUMBER:
                  (B) FILING DATE:
35
           (viii) ATTORNEY/AGENT INFORMATION:
                  (A) NAME: A. Anders Brookes
                  (B) REGISTRATION NUMBER: 36,373
                  (C) REFERENCE/DOCKET NUMBER: PZ004PCT
40
            (vi) TELECOMMUNICATION INFORMATION:
                  (A) TELEPHONE: (301) 309-8504
                  (B) TELEFAX: (301) 309-8439
45
     (2) INFORMATION FOR SEQ ID NO: 1:
            (i) SEQUENCE CHARACTERISTICS:
                  (A) LENGTH: 733 base pairs
50
                  (B) TYPE: nucleic acid
                  (C) STRANDEDNESS: double
                  (D) TOPOLOGY: linear
            (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:
55
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	GGGATCCGGA GCCCAAATCT TCTGACAAAA CTCACACATG CCCACCGTGC CCAGCACCTG	60
	AATTCGAGGG TGCACCGTCA GTCTTCCTCT TCCCCCCAAA ACCCAAGGAC ACCCTCATGA	120
5	TCTCCCGGAC TCCTGAGGTC ACATGCGTGG TGGTGGACGT AAGCCACGAA GACCCTGAGG	180
	TCAAGTTCAA CTGGTACGTG GACGGCGTGG AGGTGCATAA TGCCAAGACA AAGCCGCGGG	240
10	AGGAGCAGTA CAACAGCACG TACCGTGTGG TCAGCGTCCT CACCGTCCTG CACCAGGACT	300
	GGCTGAATGG CAAGGAGTAC AAGTGCAAGG TCTCCAACAA AGCCCTCCCA ACCCCCATCG	360
	AGAAAACCAT CTCCAAAGCC AAAGGGCAGC CCCGAGAACC ACAGGTGTAC ACCCTGCCCC	420
15	CATCCCGGGA TGAGCTGACC AAGAACCAGG TCAGCCTGAC CTGCCTGGTC AAAGGCTTCT	480
	ATCCAAGCGA CATCGCCGTG GAGTGGGAGA GCAATGGGCA GCCGGAGAAC AACTACAAGA	540
20	CCACGCCTCC CGTGCTGGAC TCCGACGGCT CCTTCTTCCT CTACAGCAAG CTCACCGTGG	600
	ACAAGAGCAG GTGGCAGCAG GGGAACGTCT TCTCATGCTC CGTGATGCAT GAGGCTCTGC	660
	ACAACCACTA CACGCAGAAG AGCCTCTCCC TGTCTCCGGG TAAATGAGTG CGACGGCCGC	720
25	GACTCTAGAG GAT	733
30	(2) INFORMATION FOR SEQ ID NO: 2:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 5 amino acids (B) TYPE: amino acid	
35	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:	
40	Trp Ser Xaa Trp Ser 1 5	
45	(2) INFORMATION FOR SEQ ID NO: 3:	
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 86 base pairs(B) TYPE: nucleic acid	
50	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:	
55	GCGCCTCGAG ATTTCCCCGA AATCTAGATT TCCCCGAAAT GATTTCCCCG AAATGATTTC	60
	CCCGAAATAT CTGCCATCTC AATTAG	86

	(2) INFORMATION FOR SEQ ID NO: 4:	
5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:	
	GCGGCAAGCT TTTTGCAAAG CCTAGGC	27
15		
	(2) INFORMATION FOR SEQ ID NO: 5:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 271 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:	
	CTCGAGATTT CCCCGAAATC TAGATTTCCC CGAAATGATT TCCCCGAAAT GATTTCCCCG	60
20	AAATATCTGC CATCTCAATT AGTCAGCAAC CATAGTCCCG CCCCTAACTC CGCCCATCCC	120
30	GCCCCTAACT CCGCCCAGTT CCGCCCCATTC TCCGCCCCAT GGCTGACTAA TTTTTTTTAT	180
	TTATGCAGAG GCCGAGGCCG CCTCGGCCTC TGAGCTATTC CAGAAGTAGT GAGGAGGCTT	240
35	TTTTGGAGGC CTAGGCTTTT GCAAAAAGCT T	271
40	(2) THEODYNETON FOR CEO TO NO. C	
40	(2) INFORMATION FOR SEQ ID NO: 6:	
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 32 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:	
50	GCGCTCGAGG GATGACAGCG ATAGAACCCC GG	32
55	(2) INFORMATION FOR SEQ ID NO: 7:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 31 base pairs(B) TYPE: nucleic acid	
60	(C) STRANDEDNESS: double	

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(D)	TOPOLOGY:	linear
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	(2) 200001 1111011	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:	
5	GCGAAGCTTC GCGACTCCCC GGATCCGCCT C	31
10	(2) INFORMATION FOR SEQ ID NO: 8:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 12 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:	
20	GGGGACTITIC CC	12
25	(2) INFORMATION FOR SEQ ID NO: 9:	
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 73 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLCGY: linear	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:	
	GCGGCCTCGA GCGGACTTTC CCGGGGACTT TCCGGGGACT TTCCATCCTG	60
40	CCATCTCAAT TAG	73
40 .		
	(2) INFORMATION FOR SEQ ID NO: 10:	
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 256 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
50	(b) TOPOLOGY: Timear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:	
٠	CTCGAGGGGA CTTTCCCGGG GACTTTCCGG GGACTTTCCA TCTGCCATCT	60
55	CAATTAGTCA GCAACCATAG TCCCGCCCCT AACTCCGCCC ATCCCGCCCC TAACTCCGCC	120
	CAGTTCCGCC CATTCTCCGC CCCATGGCTG ACTAATTTT TTTATTATG CAGAGGCCGA	180

GGCCGCCTCG GCCTCTGAGC TATTCCAGAA GTAGTGAGGA GGCTTTTTTG GAGGCCTAGG

167

CTTTTGCAAA AAGCTT 256

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(2) INFORMATION FOR SEQ ID NO: 11:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1679 base pairs(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

15 GCAGCGCACC CGGGCGATCG CTTCACGGAT GCGGACGACG TAGCCATCCT TACCTACGTG 60 AAGGAAAATG CCCGCTCGCC CAGCTCCGTC ACCGGTAACG CCTTGTGGAA AGCGATGGAG 120 20 AAGAGCTCGC TCACGCAGCA CTCGTGGCAG TCCCTGAAGG ACCGCTACCT CAAGCACCTG 180 CGGGGCCAGG AGCATAAGTA CCTGCTGGGG GACGCGCCGG TGAGCCCCTC CTCCCAGAAG 240 CTCAAGCGGA AGGCGGAGGA GGACCCGGAG GCCGCGGATA GCGGGGAACC ACAGAATAAG 300 25 AGAACTCCAG ATTTGCCTGA AGAAGAGTAT GTGAAGGAAG AAATCCAGGA GAATGAAGAA 360 GCAGTCAAAA AGATGCTTGT GGAAGCCACC CGGGAGTTTG AGGAGGTTGT GGTGGATGAG 420 30 AGCCCTCCTG ATTTTGAAAT ACATATAACT ATGTGTGATG ATGATCCACC CACACCTGAG 480 GAAGACTCAG AAACACAGCC TGATGAGGAG GAAGAAGAAG AAGAAGAAAA AGTTTCTCAA 540 CCAGAGGTGG GAGCTGCCAT TAAGATCATT CGGCAGTTAA TGGAGAAGTT TAACTTGGAT 600 35 CTATCAACAG TTACACAGGC CTTCCTAAAA AATAGTGGTG AGCTGGAGGC TACTTCCGCC 660 TTCTTAGCGT CTGGTCAGAG AGCTGATGGA TATCCCATTT GGTCCCGACA AGATGACATA 720 40 GATTTGCAAA AAGATGATGA GGATACCAGA GAGGCATTGG TCAAAAAATT TGGTGCTCAG 780 840 AAAGTCATGG TAGGTGAGGT GGTTAAAAAA AATTGTGACC AATGAACTTT AGAGAGTTCT 900 45 TGCATTGGAA CTGGCACTTA TITTCTGACC ATCGCTGCTG TTGCTCTGTG AGTCCTAGAT 960 TTTTGTAGCC AAGCAGAGTT GTAGAGGGGG ATAAAAAGAA AAGAAATTGG ATGTATTTAC 1020 50 ACCTGTCCTT GAACAAGTAT CAATGTGTTT ATGAAAGGAA GATCTAAATC AGACAGGAGT 1080 TGGTCTACAT AGTAGTAATC CATTGTTGGA ATGGAACCCT TGCTATAGTA GTGACAAAGT 1140 GAAAGGAAAT TTAGGAGGCA TAGGCCATTT CAGGCAGCAT AAGTAATCTC CTGTCCTTTG 1200 55 GCAGAAGCTC CTTTAGATTG GGATAGATTC CAAATAAAGA ATCTAGAAAT AGGAGAAGAT 1260 TTAATTATGA GGCCTTGAAC ACGGATTATC CCCAAACCCT TGTCATTTCC CCCAGTGAGC 1320 60 TCTGATTTCT AGACTGCTTT GAAAATGCTG TATTCATTTT GCTAACTTAG TATTTGGGTA 1380

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	CCCTGCTCTT	TGGCTGTTCT	TTTTTTGGAG	CCCTTCTCAG	TCAAGTCTGC	CGGATGTCTT	1440
5	TCTTTACCTA	CCCCTCAGTT	TTCCTTAAAA	CGCGCACACA	ACTCTAGAGA	GTGTTAAGAA	1500
	TAATGTTACT	TGGTTAATGT	GTTATTTATT	GAGTATTGTT	TGTGCTAAGC	ATTGTGTTAG	1560
	ATITAAAAAA	TTAGTGGATT	GACTCCACTT	TGTTGTGTTG	TTTTCATTGT	TGAAAATAAA	1620
10	TATAACTTTG	TATTCGAAAA	ААААААААА	AAAATNRCTG	CGGNCCGACA	AGGGAATTC	1679

15 (2) INFORMATION FOR SEQ ID NO: 12:

20

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1830 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

25	GCGACCGCGC CCTTCAGCTA GCTCGCTCGC TCGCTCTGCT TCCCTGCTGC CG	GGCTGCGCA (60
	TGGCTTNGGC GTTGGCGGCG CTGGCGGCGG TCGAGCNGCC TGCGSAGCCG GT	TACCAGCAG 1	20
30	TTGCAGAATG AAGAAGAGTC TGGAGAACCT GAACAGGCTG CAGGTGATGC TC	CCTCCACCT 1	80
50	TACAGCAGCA TTTCTGCAGA GAGCGCACAT NATTTTGACT ACAAGGATGA GT	rctgggttt 2	40
	CCAAAGCCCC CATCTTACAA TGTAGCTACA ACACTGCCCA GTTATGATGA AG	GCGGAGAGG 3	00
35	ACCAAGGCTG AAGCTACTAT CCCTTTGGTT CCTGGGAGAG ATGAGGATTT TC	GTGGGTCGG 3	60
	GATGATTTTG ATGATGCTGA CCAGCTGAGG ATAGGAAATG ATGGGATTTT CA	ATGTTAACT 4	20
40	TTTTTCATGG CATTCCTCTT TAACTGGATT GGGTTTTTCC TGTCTTTTTG CO	CTGACCACT 4	80
.0	TCAGCTGCAG GAAGGTATGG GGCCATTTCA GGATTTGGTC TCTCTCTAAT TA	AAATGGATC 5	40
	CTGATTGTCA GGTTTTCCAC CTATTTCCCT GGATATTTTG ATGGTCAGTA C	TGGCTCTGG 6	00
45	TGGGTGTTCC TTGTTTTAGG CTTTCTCCTG TTTCTCAGAG GATTTATCAA T	TATGCAAAA 6	60
	GTTCGGAAGA TGCCAGAAAC TTTCTCAAAT CTCCCCAGGA CCAGAGTTCT C	TTTATTTAT 7	720
50	TAAAGATGTT TTCTGGCAAA GGCCTTCCTG CATTTATGAA TTCTCTCTCA A	GAAGCAAGA	780
30	GAACACCTGC AGGAAGTGAA TCAAGATGCA GAACACAGAG GAATAATCAC C	TGCTTTAAA 8	340
	AAAATAAAGT ACTGTTGAAA AGATCATTTC TCTCTATTTG TTCCTAGGTG T	ATTTTAAAA'	900
55	ATAGTTAATG CAGAATTCTG TAATCATTGA ATCATTAGTG GTTAATGTTT G	BAAAAAGCTC S	960
	TTGCAATCAA GTCTGTGATG TATTAATAAT GCCTTATATA TTGTTTGTAG T	CATTTTAAG 1	020
60	TAGCATGAGC CATGTCCCTG TAGTCGGTAG GGGGCAGTCT TGCTTTATTC A	ATCCTCCATC 1	080

	TCAAAATGAA	CTTGGAATTA	AATATTGTAA	GATATGTATA	ATGCTGGCCA	TTTTAAAGGG	1140
	GTTTTCTCAA	AAGTTAAACT	TTTGTTATGA	CTGTGTTTTT	GCACATAATC	CATATTTGCT	1200
5	GTTCAAGTTA	ATCTAGAAAT	TTATTCAATT	CTGTATGAAC	ACCTGGAAGC	AAAATCATAG	1260
	TGCAAAAATA	CATTTAAGGT	GTGGTCAAAA	ATAAGTCTTT	AATTGGTAAA	TAATAAGCAT	1320
10	TAATTTTTTA	TAGCCTGTAT	TCACAATTCT	GCGGTACCTT	ATTGTACCTA	AGGGATTCTA	1380
10	AAGGTGTTGT	CACTGTATAA	AACAGAAAGC	ACTAGGATAC	AAATGAAGCT	TAATTACTAA	1440
	AATGTAATTC	TTGACACTCT	TTCTATAATT	AGCGTTCTTC	ACCCCACCC	CCACCCCAC	1500
15	CCCCTTATT	TTCCTTTTGT	CTCCTGGTGA	TTAGGCCAAA	GTCTGGGAGT	AAGGAGAGGA	1560
	TTAGGTACTT	AGGAGCAAAG	AAAGAAGTAG	CTTGGAACTT	TTGAGATGAT	CCCTAACATA	1620
20	CTGTACTACT	TGCTTTTACA	ATGTGTTAGC	AGAAACCAGT	GGGTTATAAT	GTAGAATGAT	1680
20	GTGCTTTCTG	CCCAAGTGGT	AATTCATCTT	GGTTTGCTAT	GTTAAAACTG	ТАААТАСААС	1740
	AGAACATTAA	TAAATATCTC	TTGTGTAGCA	CCTTTTAAAA	АААААААА	АААААААА	1800
25	ААААААААА	AANCCCGGGG	GGGGGCCCCN				1830

30 (2) INFORMATION FOR SEQ ID NO: 13:

. 35

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1212 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:.

40	TGTTTGAAGT	TGTTACTTTT	GTTTACAGCA	AAGTTTGATG	TAGTGTGCAG	TAGTGAGCTC	60
	TAGACTGATC	TTTTTCTAAA	TCAGAAAGTG	ATTAAAGTAT	GCACAACCAA	AGGCAGGTTT	120
45	TTCTTTTTCA	TTTATTCAGC	AACTATTTAT	TAAGCATCAA	CTCTGTGCCA	GGCACGTTAC	180
40	TAGCTGCTAC	ATACTGTCTG	AACATGACAT	ACGGTTAAGT	AACTTTACAA	TTATTATCAA	240
	ATACTTCAAT	GTAGATATTT	CTTAAGTTGA	AATAGCATTA	ACTAGGATAA	TGCTTTCATG	300
50	TTATTTTATT	TGTCTTGTGA	TAGAAATTCA	ACTTTGTACC	ATCTTAAAAC	TAGGTTGCTA	360
	TAAAAATAGG	AGGATGAAGT	CAATAAAGTT	TATGCCAGTT	TAAAAACTGG	AAGGAAAAGG	420
55	TAAGAGCTCT	CCATTATAAA	ATAGTTGCAT	TCGGTTAATT	TTTACACATT	AGTGCATTGC	480
33	GTATATCAAC	TGGCCCTCAA	TGAAGCATTT	AAGTGCTTGG	AATTTTACTA	AACTGACTTT	540
	TTTGCAACTT	TGGGAGATTT	TTGAGGGGAG	TGTTGAAAAT	TGCCAAACAC	TCACCTCTTA	600
60	CTCAAAACTT	CAAATAAAAT	ACACATTTTC	AAGAGGGAGC	ACCTTTTATA	TTTGATAAGT	660

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	TTTCATTATA	AACCTTATAA	TACCAGTCAC	AAAGAGGTTG	TCTGTCTATG	GTTTAGCAAA	720
5	CATTTGCTTT	TCTTTTTGGA	AGTGTGATTG	CAATTGCAGA	ACAGAAAGTG	AGAAAACACT	780
J	GCCAGCGGTG	ATTGCTACTT	GAGGTAGTTT	TTTACAACTA	CCATTTCCCC	TCCATGAAAT	840
	TATGTGAAAT	TTATTTTATC	TTTGGGAAAA	GTTGAGAAGA	TAGTAAAAGA	ATTAGGAATT	900
10	TAAAATTACA	GGGAAAAATA	TGTAAGTGAA	AAGCAATAAA	TATTTTGTTC	ACTITGCTAT	960
	CAAGATGTTC	ACTATCAGAT	ATTTATTATA	TGGCAGCAAT	TTATATTTT	AATCATTGCC	1020
15	CATTAATAGA	CGCAGTAAAA	TATTTTTGAA	TCAGACATTT	GGGGTTTGTA	TGTGCATTAA	1080
13	AATTGTCTTT	TGTACTGTAA	GTTACTGTTA	ATTTGAATAT	TTTATTGAAC	TGTCTCCCTG	1140
	TGCCTTTATA	ATATAAAGTT	GTTTCTACAA	CTTTTAATGA	TCTTAATAAA	GAATACTTTA	1200
20	AGAAAAAAA	AA	·				1212

25 (2) INFORMATION FOR SEQ ID NO: 14:

30

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2061 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:

35	GGTTTTCCTC	CGACTTCCGG	ACATCTCCCT	GGGAGTCGCG	CAGAGTGGAG	TCAAAGGCAA	60
	CCAGTGCTCG	CTGCGGTCTC	TGGGGATCGG	GACCGCGGCG	GCGGCCCGCG	AGCGGGATGT	120
40	TCCGGGGCTT	GAGCAGTTGG	TTGGGCTTGC	AGCAGCCGGT	GGCAGGCGGT	GGGCAGCCCA	180
10	ATGGAGATGC	TCCACCCGAG	CAGCCGTCCG	AGACGGTGGC	TGAGTCTGCG	GAGGAGGAGC	240
	TGCAGCAAGC	GGGAGACCAG	GAGCTCCTCC	ACCAGGCCAA	AGACTTCGGC	AACTATTTAT	300
45	TTAACTTTGC	ATCTGCTGCC	ACAAAAAAGA	TAACTGAATC	AGTTGCTGAA	ACAGCACAAA	360
	CAATAAAGAA	ATCCGTAGAA	GAAGGAAAAA	TAGATGGCAT	CATTGACAAG	ACAATTATAG	420
50	GAGATTTTCA	GAAGGAACAG	AAAAATTTG	TTGAAGAGCA	ACATACAAAG	AAGTCAGAAG	480
50	CAGCTGTGCC	CCCATGGGTT	GACACTAACG	ATGAAGAAAC	AATTCAACAA	CAAATTTTGG	540
	CCTTATCAGC	TGACAAGAGG	AATTTCCTTC	GTGACCCTCC	GGCTGGCGTG	CAATTTAATT	600
55	TCGACTTTGA	TCAGATGTAC	CCCGTGGCCC	TGGTCATGCT	CCAGGAGGAT	GAGCTGCTAR	660
	CAAGATGAGA	TTTGCCCTCG	TTCCTAAACT	TGTGAAGGAA	GAAGTGTTCT	GGAGGAACTA	720
60	CTTTTACCGC	GTCTCCCTGA	TTAAGCAGTC	AGCCCAGCTC	ACGGCCCTGG	CTGCCCAACA	780

GCAGGCCGCA GGGAAGGGA GAGAAGAGCA ATGGCAGAGA GCAAGATTTG CCGCTGGAGA

	GGCAGTACGG CCCAAAACGC CACCCGTTGT AATCAAATCT CAGCTTAAAA CTCAAGAGGA	900
5	TGAGGAAGAA ATTTCTACTA GCCCAGGTGT TTCTGAGTTT GTCAGTGATG CCTTCGATGC	960
	CTGTAACCTA AATCAGGAAG ATCTAAGGAA AGAAATGGAG CAACTAGTGC TTGACAAAAA	1020
10	GCAAGAGGAG ACAGCCGTAC TGGAAGAGGA TTCTGCAGAT TGGGAAAAAG AACTGCAGCA	1080
10	GGAACTTCAA GAATATGAAG TGGTGACAGA ATCTGAAAAA CGAGATGAAA ACTGGGATAA	1140
	GGAAATAGAG AAAATGCTTC AAGAGGAAAA TTAGCTGTTC CTGAAATAGA AGAATAATCC	1200
15	TTAACAGTCT GCAAACTGAC ATTAAATTCT AGATGTTGAC AATTACTGAA TCAGAAGGCA	1260
	TGAAAGAGTA TAATITTATG AAATTCAAAA TTATTCTTTT TTCAAGTTGA AACTTGCCTC	1320
20	TTCTACTITA AAAAAGTATA TAGAACAGTT ACTTCTAATA ATCAGAAAGA GATGTTTTAT	1380
	AGAACATTTC TTTAATATAA AGTTAGAGAT GTCTTCATAG GCAGTATGGC TATCTTTGCC	1440
	ACAGAAACAT AAGTAAAATT TTAGAGTTCT GTTTTCCATG AGGTCAAAAA TATAATTTAT	1500
25	TCCTCAGTCA TGGTTTTCTA AATATCTGTA CTCCACATTC CATTTTAATT GATATGAGGG	1560
	TGTTAAAGTA CCTACTTAAT GGGTTGATTA CTATCAAAAT GACCAAATTA TACCAAAGAA	1620
30	CTTAAGAGGA AGCACTTTCA GAACTATTCA CTTGCCAGGT ATTTTCTAAA ATTCCACCTG	1680
	AAAGCCAAAA GATAAAATAC ATNAGTTGGA TTTTAATGAT ATAAGCATCA CACAATTTTA	1740
	CATTAAGAAA TACTGTGCAG CCCATGCGTG GTGGCTCAGG CCTGTAATCC CAGCANTTTG	1800
35	GGAGGCCGAG GTGGGCAGAT CACCGGAGGT CAGGAGTTCG AGACCAGCCT TGCCAACATA	1860
	GTGAAACCCT GTCTTTACTA AAAATACAAA AATTAGCCGG GCATGGTGGC AGGCACCTGT	1920
1 0	AATCCCAGCT ACTAGGGAGG CTTTTGAACC CAGGAGGCAG AGGTTGCAGC GAGCTGAGAT	1980
	CGCGCCACTG CACTCCAGCC TGGGTGATAG AGTGAGATTC AGTCTCAAAA AAAAAAAAA	2040
	AAAAAAAA AATGACCTCG A	2061
15		
	(2) INFORMATION FOR SEQ ID NO: 15:	
50	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1412 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:	
	CCCTTCATCT GCGTTGCCAG GAACCCTGTC AGCAGAAACT TCTCAAGCCC CATCCTTGCC	60
50	AGGAAGCTCT GTGAAGGTGC TGCTGATGAC CCAGATTCCT CCATGGTCCT CCTGTGTCTC	120
	constitution	140

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	CTGTTGGTGC	CCCTCCTGCT	CAGTCTCTTT	GTACTGGGGC	TATTTCTTTG	GTTTCTGAAG	180
5	AGAGAGAGAC	AAGAAGAGTA	CATTGAAGAG	AAGAAGAGAG	TGGACATTTG	TCGGGAAACT	240
5	CCTAACATAT	GCCCCCATTC	TGGAGAGAAC	ACAGAGTACG	ACACAATCCC	TCACACTAAT	300
	AGAACAATCC	TAAAGGAAGA	TCCAGCAAAT	ACGGTTTACT	CCACTGTGGA	AATACCGAAA	360
10	AAGATGGAAA	ATCCCCACTC	ACTGCTCACG	ATGCCAGACA	CACCAAGGCT	ATTTGCCTAT	420
	GAGAATGTTA	TCTAGACAGC	AGTGCACTCC	CCTAAGTCTC	TGCTCAAAAA	AAAAACAATT	480
	CTCGGCCCAA	AGAAAACAAT	CAGAAGAATT	CACTGATTTG	ACTAGAAACA	TCAAGGAAGA	540
15	ATGAAGAACG	TTGACTTTTT	TCCAGGATAA	ATTATCTCTG	ATGCTTCTTT	AGATTTAAGA	600
	GTTCATAATT	CCATCCACTG	CTGAGAAATC	TCCTCAAACC	CAGAAGGTTT	AATCACTTCA	660
20	TCCCAAAAAT	GGGATTGTGA	ATGTCAGCAA	ACCATAAAAA	AAGTGCTTAG	AAGTATTCCT	720
	ATAAAAATGT	AAATGCAAGG	TCACACATAT	TAATGACAGC	CTGTTGTATT	AATGATGGCT	780
25	CCAGGTCAGT	GTCTGGAGTT	TCATTCCATC	CCAGGGCTTG	GATGTCAGGA	TTATACCAAG	840
25	AGTCTTGCTA	CCAGGAGGC	AAGAAGACCA	AAACAGACAG	ACAAGTCCAG	CAGAAGCAGA	900
	TGCACCTGAC	AAAAATGGAT	GTATTAATTG	GCTCTATAAA	CTATGTGCCC	AGCAYTATGC	960
30	TGAGCTTACA	CTAATTGGTC	AGACATGCTG	TCTGCCCTCA	TGAAATTGGC	TCCAAATGAW	1020
	TGAACTACTT	TCATGAGCAG	TTGTAGCAGG	CCTGACCACA	GATTCCCAGA	GGGCCAGGTG	1080
2.5	TGGATCCACA	. GGACTTGAAG	GTCAAAGTTC	: ACAAAGATGA	AGAATCAGGG	TAGCTGACCA	1140
35	TGTTTGGCAG	ATACTATAAT	GGAGACACAG	AAGTGTGCAT	· GGCCCAAGGA	CAAGGACCTC	1200
	CAGCCAGGCT	TCATTTATGC	ACTTGTCTGC	: AAAAGAAAAG	TCTAGGTTT	AAGGCTGTGC	1260
40	CAGAACCCAT	CCCAATAAAG	AGACCGAGTO	TGAAGTCACA	TTGTAAATCT	AGTGTAGGAG	1320
	ACTTGGAGTO	: AGGCAGTGAG	ACTGGTGGG	CACGGGGGG	ANTGGGTAM	GTAAACCTTT	1380
	TAAAGATGGT	TAATTCNTCA	TTAGTGTTT	TT			1412
45							

(2) INFORMATION FOR SEQ ID NO: 16:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1052 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

55 (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

TTCCTCTCCT CTCTCTACCC CTCCTGTCTC TCCTCCCCTC CTCTCTCTTC CTCTCCTCTC 60

50

	TCTCTTCCTC	TCCTCTCTCT	TCCCTTCCTG	TCTCTCTTCC	CCTCCTCTCT	CTCTTCCTGT	120
	CCTCTATCTC	TTCCCCTCCT	CTATCTCTTC	CTCTCCTCTC	TCTCTTCCTC	TCCTCTCTCT	180
5	CTCTTSCTTT	CTTCTCTCTC	TCCTGTCTCG	GCTGTTGTGG	GTTGCAGGTT	GGGTGCTGCT	240
	GTTGTGGTCC	TTCCCAGAAA	CTGCCAGTAG	AGGGCAGCCT	GGGCATCCTA	ATGCTTACTC	300
10.	TGGTTGTTAC	ACAAAGAAAA	TATTGGGGTC	ACTGGCGAGC	CCACCCACAC	TCACCAGAAT	360
10,	CTCCACTGTA	GTCCCCTAA	CAAACAGCCC	TTCACTTCCT	CTCCCACTTC	AGCAATTTGT	420
	ATTTTGATGC	CATTGGCCTC	AGATCAGAGT	GTTTTAAATC	ATCACGCCCT	GGCTTATCCC	480
15	TGGTCGAGCC	AGGACACGGG	GTGCTTCAGT	GGTCTGTCA	CCCTCTCTCC	TTGAAGCATG	540
	TTGCTTTTAT	TTATTTACTT	TTACTCTCAC	CCTGCTCCTG	TACCAGCAGG	GGCCACTTCA	600
20	AAGCCAAGGT	ACAGGGTGAT	AACTTGTGGT	CCAGCATCAG	TTTTCTCCAC	TTCTTTCTCC	660
20	CACTCACCCC	CAGCAAGGTG	CCTGGGGAGA	CTTGAGCAGA	TGTTTCATTT	TGGCCTGGCC	720
	AGTGGCTGAA	AGCAGGCCTC	CAATGCACTG	TGACCTCTGG	CTTCCCCAGC	AGCTTTCCCA	780
25	GAGAGGCAGA	GGGCCTTCC	ACAGCCCGGG	TTCTCCTGCT	GCCTCCTGCC	TGCTGCAGCT	840
	GCAGGCATTC	TGAGGGGCAA	CGTGGAGGAA	GGGCCAGGGA	TGCATGGGAT	TTTAATTGTT	900
30	TCATCACACC	TTCCCCGTGG	CAAAGAAACA	GTCAGTCCTC	TTCAGGTGTC	TTCTGGATTT	960
	CTGGTGATGG	ACAGAGAAAT	CTTTTTACAG	TTTCAAATTA	TGTTCAACAA	ATAAAAATTG	1020
	CATTTTTTAT	TTTGGAAAAA	АААААААА	AA			1052
35							
	(2) INFORM	ATION FOR S	EO ID NO: 1	7 :			,
40	(i)	SEQUENCE C	HARACTERIST	ics:			
		(A) LEN	NGTH: 683 ba	se pairs			
		(C) STF	CANDEDNESS:	double			
45	(xi) SEQUENCE): 17:		
						TTAGACAAAG	60
50						GCTGATAGCT	120
		-				TTATTCAGAA	180
						GTCTTTCTGT	240
55						ATTTCTTT	300
						GTGAGCTCTA	360
60						AAATAACATT	420

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	ATCATATGTW ATTGGCATAT AAATTACAGA TGTWTCTATG ACTAAAAACC CTGTGGATAT	480
5	WAACCMAATG CAGATAAWTW TAATAAAATW TWTAAAAATW TWATCMAATA ATGATAGTGC	540
5	TATTCAAATA CTTCAAATTT GCACAGTGAT TTATTTCTTA AAATATGTTA ACACATGTGA	600
	GCCAATACAC TGAGGTCACT GGATAAATAA ACAGATTCTT GCAAAAAAAA AAAAAAAAAA	660
0	ACTCGAGGGG GGCCCGTACC CTT	683
15	(2) INFORMATION FOR SEQ ID NO: 18:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1054 base pairs (B) TYPE: nucleic acid	
20	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:	
25	AAACTCATTT AGGTGACACT ATAGAAGGTA CGCCTGCAGG TACCGGTCCG GAATTCCCGG	60
	GTCGACCCAC GMGNCCGGCG ACAAGATGGC AGCAGCGTGT CGGAGCGTGA AGGGCCTGGT	120
•	GGCGGTAATA ACCGGAGGAG CCTCGGGCCT GGGCCTGGCC ACGGCGGACG ACTTGTGGGG	180
30	CAGGGAGCCT CTGCTGTGCT TCTGGACCTG CCCAACTCGG GTGGGGAGGC CCAAGCCAAG	240
	AAGTTAGGAA ACAACTGCGT TTTCGCCCCA GCCGACGTGA CCTCTGAGAA GGATGTGCAA	300
35	ACAGCTCTGG CTCTAGCAAA AGGAAAGTTT GGCCGTGTGG ATGTAGCTGT CAACTGTGCA	360
	GGCATCGCGG TGGCTAGCAA GACGTACAAC TTAAAGAAGG GCCAGACCCA TACCTTGGAA	420
40	GACTTCCAGC GAGTTCTTGA TGTGAATCTC ATGGGCACCT TCAATGTGAT CCGCCTGGTG	480
40	GCTGGTGAGA TGGGCCAGAA TGAACCAGAC CAGGGAGGCC AACGTGGGGT CATCATCAAC	540
	ACTGCCAGTG TGGCTGCCTT CGAGGGTCAG GTTGGACAAG CTGCATACTC TGCTTCCAAG	600
45	GGGGGAATAG TGGGCATGAC ACTGCCCATT GCTCGGGATC TGGCTCCCAT AGGTATCCGG	660
	GTGATGACCA TTGCCCCAGG TCTGTTTGGC ACCCCACTGC TGACCAGCCT CCCAGAGAAA	720
50	GTGTGCAACT TCTTGGCCAG CCAAGTGCCC TTCCCTAGCC GACTGGGTGA CCCTGCTGAG	78
50	TATGCTCACC TCGTACAGGC CATCATCGAG AACCCATTCC TCAATGGAGA GGTCATCCGG	84
	CTGGAŢGGGG CCATTCGTAT GCAGCCTTGA AGGGAGAAGG CAGAGAAAAC ACACGCTCCT	90
55	CTGCCCTTCC TTTCCCTGGG GTACTACTCT CCAGCTTGGG AGGAAGCCCA GTAGCCATTT	96

TGTAACTGCC TACCAGTCGC CCTCTGTGCC TAATAAAGTC TCTTTTTCTC ACANAAAAAA

ΑΑΑΑ ΑΑΑΑΑΑΑΑ ΑΚΑΑΑΑΑΑΑ

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1020 1054

(2) IN	FORMATION	FOR	SEQ	ID NO:	19:
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(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1393 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 19:

	•	
15	GGAACAAGCT GGGATATGTG AGCGTTAAGC TACTCACATC CTTCAAAAAG GTGAAACATC	60
• 5	TTACACGGGA CTGGAGAACC ACAGCACATG CTTTGAAGTA TTCAGTGGTC CTTGAGTTGA	120
	ATGAGGNCCA CCGGAAGGTG AGGAGGACCA CCCCCGTCCC ACTGTTCCCC AACGAGAACC	180
20	TCCCCAGCAA GATGCTCCTG GTCTATGATC TCTACTTGTY TCCTAAGCTG TGGGCTCTGG	240
	CCACCCCCA GAAGAATGGG AAGGGTGCAA GARAAGGTGA TGGAACACCT GCTCAAGCTT	300
25	TTTGGGACTT TTGGAGTCAT CTCATCAGTG CGGATCCTCA AACCTGGGAG AGAGCTGCCC	360
23	CCTGACATCC GGAGGNTCCA GCAGCCGCTA CAGCTCCTCT GACCCCGAGA GCAACCCCAC	420
	ATCCCCTATG GCGGGCCGAC GGCACGNGKC CACCAACAAG CTCAGCCCGT CTGGCCACCA	480
30	GAATCTCTTT CTGAGTCCAA ATGCCTCCCC GTGCACAAGT CCTTGGAGCA GCCCCTTGGC	540
	CCAACGCAAA GGCGTTTCCA GAAAGTCCCC ACTGGCGGAG GAAGGTAGAC TGAACTGCAG	600
35	CACCAGCCCT GAGATCTTCC GCAAGTGTAT GGATTATTCC TCTGACAGCA GCGTCACTCC	660
33	CTCTGGCAGC CCCTGGGTCC GGAGGCGTCG CCAAGCCGAG ATGGGGACCC AGGAGAAAAG	720
	CCCCGGTACG AGTCCCCTGC TCTCCCGGAA GATGCAGACT GCAGATGGGS TACCCGTAGG	780
40	TNGCTTGAGG TTGCCCAGGG GTCCTGACAA CACCAGAGGA TTTCATGGCC ATGAGAGGAG	840
	CAGGGCCTGT GTATAAATAC CTTCTATTTT TAATACAAGC TCCACTGAAA ACCACCTTCG	900
15	TTTTCAAGGT TCTGACAAAC ACCTGGCATG ACAGAATGGA ATTCGTTCCC CTTTGAGAGA	960
45	TTTTTTATTC ATGTAGACCT CTTAATTTAT CTATCTGTAA TATACATAAA TCGGTACGCC	1020
	ATGGTTTGAA GACCACCTTC TAGTTCAGGA CTCCTGTTCT TCCCAGCATG GCCACTATTT	1080
50	TGATGATGGC TGATGTGTGT GAGTGTGATG GCCCTGAAGG GCTGTAGGAC GGAGGTTCCC	1140
	TGGGGGAAGT CTGTTCTTTG GTATGGAATT TTTCTCTCTT CTTTGGTATG GAATTTTTCC	1200
. .	CTTCAGTGAC TGAGCTGTCC TCGATAGGCC ATGCAAGGGC TTCCTGAGAG TTCAGGAAAG	1260
55	TTCTCTTGTG CAACAGCAAG TAGCTAAGCC TATAGCATGG TGTCTTGTAG GACCAAATCG	1320
	ATGITACCTG TCAAGTAAAT AAATAATAAA ACACCCAACT GGGAGTGCTG AAAAAAAANA	1380
60	ANNAAAAAC TCG	1393

5	101	TATE OF TAKE	EOB	CEO	TD	NO	20
5	(z)	INFORMATION	FUR	SEQ	ΤU	MO:	20:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1215 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

15	AGGAAAAGTT	TTCCNAATTG	GAAAGCGGGC	AGTGAGCGCA	ACGCAATTAA	TGTGAGTTAG	60
	NTCANTCATT	AGGCACCCCA	GGCTTTACAC	TTTATGCTTC	CGGNTCGTAT	GTTGTGTGGA	120
20	ATTGTGAGCG	GATAACAATT	TCACACAGGA	AACAGCTATG	ACCATGATTA	CGCCAAGCTN	180
20	TAATACGACT	CACTATAGGG	AAAGCTGGTA	CGCCTGCAGG	TACCGGTCCG	GAATTCCCGG	240
	GTCGACCCAC	GCGTCCGCCC	ACGCGTCCGT	GAAAATCCGA	AGTGCCGCGG	AAAGTGGAGG	300
25	TGAGGGCCGC	CCGCCCTAGA	GGTGCCCGTC	CGAGAGGCAG	AGCTGACAAG	GAAGGTTTCG	360
	AGCGTTTTGC	TGGCAAAGGG	ATTTCTTACA	ACCTCCAGGC	ATGCGTCTTT	CTGCCCTGCT	420
30	GGCCTTGGCA	TCCAAGGTCA	CTCTGCCCCC	CCATTACCGC	TATGGGATGA	GCCCCCAGG	480
30	CTCTGTTGCA	GACAAGAGGA	AGAACCCCCC	ATGGATCAGG	CGGCGCCCAG	TGGTTGTGGA	540
	ACCCATCTCT	GATGAAGACT	GGTATCTGTT	CTGTGGGGAC	ACGGTGGAGA	TCCTAGAAGG	600
35	CAAGGATGCC	GGGAAGCAGG	GCAAAGTGGT	TCAAGTTATC	CGGCAGCGAA	ACTGGGTGGT	660
	CGTGGGAGGG	CTGAACACAC	ATTACCGCTA	CATTGGCAAG	ACCATGGATT	ACCGGGGAAC	720
40	CATGATCCCT	AGTGAAGCCC	CCTTGCTCCA	CCGCCAGGTC	AAACTTGTGG	ATCCTATGGA	780
40	CAGGAAACCC	ACTGAGATCG	AGTGGAGATT	TACTGAAGCA	GGAGAGCGGG	TACGAGTCTC	840
	CACACGATCA	GGGAGAATTA	TCCCTAAACC	CGAATTTCCC	AGAGCTGATG	GCATCGTCCC	900
45	TGAAACGTGG	ATTGATGGCC	CCAAAGACAC	ATCAGTGGAA	GATGCTTTAG	AAAGAACCTA	960
	TGTGCCCTGT	CTAAAGACAC	TGCAGGAGGA	GGTGATGGAG	GCCATGGGGA	TCAAGGAGAC	1020
50	CCGGAAATAC	AAGAAGGTCT	ATTGGTATTG	AGCCTGGGGC	AGAGCAGCTC	CTCCCCAACT	1080
30	TCTGTCCCAG	CCTTGAAGGC	TGAGGCACTT	CTTTTTCAGA	TGCCAATAAA	GAGCACTTTA	1140
	TGAGTCCTCC	ААААААААА	АААААААА	AAAAAAAAAA	. AAAAAAAAA	AAAAAAAAA	1200
55	AAAAGGGGCG	GCCGC					1215

^{60 (2)} INFORMATION FOR SEQ ID NO: 21:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2042 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

				. 550 15 140	. 41.		
10	CTGCATCCAG	GCGCAGAATA	ACCTGGGTAT	CTTGTGGTCT	GAAAGAGAGA	AATTGAAACT	60
	GCACAGGCTT	ACCTAGAGTC	ATCAGAAGCA	СТАТАТААТС	AGTATATGAA	AGAGGTTGGG	120
15	AGTCCTCCTC	TTGATCCTAC	TGAGCGTTTT	CTTCTGAAGA	AGAGAAACTT	ACTGAACAAG	180
13	AGAGATCAAA	AAGATTTGAA	AAGGTTTATA	CTCATAACCT	ATATTACCTA	GCTCAAGTCT	240
	ACCAGCATCT	GGAAATGTTT	GAGAAGGCTG	CTCACTATTG	CCATAGTACA	CTAAAACGCC	300
20	AGCTTGAGCA	CAATGCCTAC	CATCCTATAG	AGTGGGCTAT	CAATGCTGCT	ACCTTGTCAC	360
	AGTTTTACAT	CAATAAGCTA	TGCTTTATGG	AGGCCAGGCA	CTGTTTATCA	GCTGCTAATG	420
25	TCATTTTTGG	TCAAACTGGA	AAGATCTCAG	CCACAGAAGA	CACTCCTGAA	GCTGAAGGAG	480
	AAGTGCCAGA	GCTTTATCAT	CAAAGAAAGG	GGGAAATAGC	AAGGTGCTGG	ATCAAATACT	540
	GTTTGACTCT	CATGCAGAAT	GCCCAACTCT	CCATGCAGGA	CAACATAGGA	GAGCTTGATC	600
30	TTGATAAACA	GTCTGAACTT	AGAGCTTTAA	GGAAAAAAGA	ACTAGATGAG	GAGGAAAGCA	660
	TTCGGAAAAA	AGCTGTGCAG	TTTGGAACCG	GTGAACTGTG	TGATGCCATC	TCTGCAGTAG	720
35	AAGAGAAAGT	GAGCTACTTG	AGACCTTTAG	ATTTTGAAGA	AGCCAGAGAA	CTTTTCTTAT	780
	TGGGTCAGCA	CTATGTCTTT	GAGGCAAAAG	AGTTCTTTCA	GATTGATGGT	TATGTCACTG	840
	ACCATATTGA	AGTTGTCCAA	GACCACAGTG	CTCTGTTTAA	GGTGCTTGCA	TTCTTTGAAA	900
40	CTGACATGGA	GAGACGGTGC	AAGATGCATA	AACGCRGAAT	AGCCATGCTA	GAGCCCCTAA	960
	CTGTAGACCT	GAATCCACAG	TATTATCTGT	TGGTCAACAG	ACAGATCCAG	TTTGAAATTG	1020
45	CACATGCTTA	CTATGATATG	ATGGATTTGA	AGGTTGCCAT	TGCTGACAGG	CTAAGGGATC	1080
	CTGATTCACA	CATTGTAAAA	ATAAATA	ATCTTAATAA	GTCAGCACTG	AAGTACTACC	1140
	AGCTCTTCTT	AGACTCCCTG	AGAGACCCAA	ATAAAGTATT	CCCTGAGCAT	ATAGGGGAAG	1200
50	ATGTTCTTCG	CCCTGCCATG	TTAGCTAAGT	TTCGAGTTGC	CCGTCTCTAT	GGCAAAATCA	1260
	TTACTGCAGA	TCCCAAGAAA	GAGCTGGAAA	ATTTGGCAAC	ATCATTGGGA	ACATTACAAA	1320
55	TTTATTGTTG	ATTACTGTGA	AAAGCATCCT	GAGGCCGCCC	AGGAAATAGA	AGTTGAGCTA	1380
50	GAACTTAGTA	AAGAGATGGT	TAGTCTTCTC	ССААСААААА	TGGAGAGATT	CAGAACCAAG	1440
	ATGGCCCTGA	CTTAATCCTT	GTTTTTAAAG	AAAGGAAATG	TGCAATATTG	AAGTGATCTT	1500
60	TTTCCCTAGT	CAGACAGGCC	CAATTCCATT	GTGATGTTTA	CCTTTATAGC	CAGGTGAGTG	1560

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	CAGTTTGAAC TTGAGATACA GTCAACTGAG TGTTTGCTAG GATCCTAAGG AACATAAAGT	1620
5	TAATTAAAAA CTTACACCTA ATTATGTAAA TTGCCTTGTT AAAGACATGT GATTTGTATT	1680
J	TTAGATGCTT GTTTCCTATT AAAATACAGA CATTTCTACC CTCAGTTTCT AAATGTAGAC	1740
	TATTTGTTGG CTAGTACTTG ATAGATTCCT TGTAAGAAAA AATGCTGGGT AATGTACCTG	1800
10	GTAACAAGCC TGTTAATATA TTAAGATTGA AAAAGTAACT TCTATAGTTA CTCCTTCTAA	1860
	AATATTTGAC TTCCTACATT CCCCCCACCC AAAATCTTTC CCTTTTGAAA ATACTAAAAA	1920
15	CTAAGTTATG TTATTATAAA GTGTAAAATG GTTTGTCTTA ATTATAGGAG AAAAAGGCCT	1980
13	TGTTAGAAAT AAAATAAACT GACTTATTTC ACTAATGAAA AAAAAAAAA AAAAAAAAA	2040
	TT	2042
20		
	(2) INFORMATION FOR SEQ ID NO: 22:	
25		
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1872 base pairs	
	(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
30	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:	
	GGGTCGACCC ACGCGTCCGA TTGGCCTAGA GCTCCTGTGA CCGAGAGCGC CACGGAAGCC	60
35	TGGGGATGAT GTCGGGCAGC TTTATTCTTT GCTTGGCTTT GGTAACTAGG TGGTCCCCTC	120
	AAGCATCCTC AGTTCCTCTT GCTGTTIATG AATCTAAGAC AAGGAAGTCC TATAGAAGCC	180
40	AAAGGGACAG GGACGGAAAG GACAGGTCCC AAGGGATGGG GCTGTCTTTA CTTGTGGAAA	240
	CCAGGAAATT GCTCCTCTCA GCCAACCAAG GTTGACCACA CACCACCCTT CCGGAGCAGC	300
	TCAGTCAGCC CTCGGGGACG RGAAACCACA AGCGCAGAGA CGCTGAGGCC CAGGCAGGTG	360
45	AAGAGGAAGT GGCTTTGGGT TTTTAAAGTA GGTGAGCGTG ACCTCTCTGA CTGCTTCTTC	420
	CCCGGGGGGG ACTGCAAACC GCTCAGGGTT GCGGCAGAGC CATGGACTTC CGGTCCCTGC	480
50	AACGGGTGAC CTAAGCGTGG TGCACCCATC AGTCACGCAG GAGGACTGAC TTGACAGACG	540
	AAAGACAAGC CCGGATGACA CAGGGTGAGA AGAGTCAGGG CCGCACCTCT GTCCCTGCAA	600
	ACCAACAGGT GCATGGTGAG TGTGGCAGTC CCCACAGCTC CACAATGGGC TCCCCCGCCA	660
55	ACGGGGACGA CAGGGATCTT CAGGAACTTC TGACCTCACC AAGTCAAGTG GACCACTCTC	720
	CACTCCACGA GGATGTGAAA CGGTTCTTTA AAATGGGATT TTAGAGCCTC GGGAATGCAT	780
	CTCCCTCCA TCTTTCATAT TATCCCTCAC CATACATTCA TTTCTTCCAA CATACTCCAA	840

	AAGATATAAG	CTGCAGTAAT	TTGCTCTTTG	AATGACCGTC	ACCCCCAGTA	TAGGATATGC	900
	TTGTATCCCC	CCGTCACTCC	TCCGCCTGTT	TTTTAAACTT	TTCCACCACC	TGCGTCCAAA	960
5	AAGAATGTTA	TAGCGAGTGC	TCTTAAATGT	TGAACCTGGG	TGTTGCTTCC	GGGCCAGTCT	1020
	GCGTGGCŤCC	ATGAAAAGCT	CACTGCTGCC	CCAGCCGGGC	TTCTTAGAGG	AGGTCAGTTG	1080
10	TCCTATGTAT	CATCATTTAC	TCTGGGAATC	CTACTGTGAA	ATCATGTCTG	TATTTTTCTG	1140
10	GAGCAGTTCA	CATAGAGTAG	AATGTGGAAT	TTCCCGTGAA	CGTCTCCTTC	CTCCCCGTA	1200
	TCTGCCGCCT	GTCACTTCGC	CACCGTGCTA	GAATACTGTT	GTGTTGTAAG	ATGACTAATT	, 1260
15	TTAAAAGAAC	CTGCCCTGAA	AAGTTCTTAG	AAACGCAATG	AAAGGGAGGA	ACTTGTCCTT	1320
•	TACCCAGTTT	TTCCTTTGTA	GGATGGGAAA	GTATAAAAAG	GCACAGAAGG	TTGTCATGGG	1380
20	CTGTTCCTTG	GGGGTTTTTA	TCCTGCTCAC	CGTGGAGATA	AGCCTGCGGC	TTGTCTAACC	1440
20	AGCGCAGCGM	AAAGGTCTCA	ATGCCTTTTG	GTAACATCCG	TCATTGCAGA	AGAAAGTTTA	1500
	CACGACGTCA	AAAAGTGACG	TTCATGCTAA	GTGTTTTTCC	AGAAATATTG	GTTTCATGTT	1560
25	TCTTATTKGC	TCTGCCTCCT	GTGCTTATAT	CATCCAAAAA	CTTTTTAAAA	AGGTCCAGAA	1620
	TTCTATTTTA	ACCTGATGTT	GAGCACCTTT	AAAACGTTCG	TATGTGTGTT	GCACTAATTC	1680
30	TAAACTTTGG	AGGCATTTTG	CTGTGTGAGG	CCGATCGCCA	CTGTAAAGGT	CCTAGAGTTG	1740
50	CCTGTTTGTC	TCTGGAGATG	GAATTAAACC	AAATAAAGAG	CTTCCACTGG	AGGCTTGTAT	1800
	TGACCTTGTA	ACTATATGTT	AATCTCGTGT	AAAATAAAT	TATAACTTGT	GAAAAAAAA	1860
35	AAAAAAAAAC	MT					1872

40 (2) INFORMATION FOR SEQ ID NO: 23:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 289 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

50 CATTTACCCA CCTATCAACA TGTTTGCTTT CTCTTTTGTT GGTGAGAATG AGTGGCTTCT 60

TGCTCCTAGC TAGAGCCAGT CCTTCCATAT GTGCTTTAGA TTCTTCCTGT TTTGTTCAAG 120

AATATTGCTC AAGCTATTCT TCCTCCTGTT TCCTGCATCA GCATTTCCCC TCTCTACTAG 180

ATCATCTCTG TCAGTAAATG AACATGTTGT TGTTTCTCCT AGAAGTACTG TTTCTATATC 240

TAGATAGTAC TCTAGCTAGA GTTAAAAAAA AAAAAAAAA CCTNGGGGG 289

(2) INFORMATION FOR SEQ ID NO: 24:

5	(i)	SEQUENCE CHARACTERISTICS:	
		(A) LENGTH: 3533 base pairs	
		(B) TYPE: nucleic acid	
		(C) STRANDEDNESS: double	
		(D) TOPOLCCY, linear	

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:

	TTTTATTTAC	TTCAAATTAA	CTGTACTTTA	CTCAAATAGA	AAANGAATAA	TTTTCACATT	60
15	ATGAAGCTAC	ACAATTCCAA	AATACACATG	CTGAGGCTCT	TTTTAAGTCC	GAATTGTCTA	120
	GTAATTACAA	AAAAGTGAAG	AGTTTACAGA	TATACAAGGA	AATAAAGGCG	AATTATTGCA	180
20	AAGAAAACAA	GTTTAATTTC	ACTTTGAATG	ACAACGATTT	TTCTGGAAAG	CAGATACTIC	240
20	ACTCCTTTAA	GTTTCCACCC	AAGCCACAAT	AATTTCAAAC	GGTCTTGCGG	ATGACCCAGC	300
	TGGTCACTCT	TGTTTATGTG	GGGACTGGAG	GTAATGAGAG	CCAAAAAAAG	TGCTATAAAC	360
25	CTAATTTGGC	TAGAGCAAGT	TCACACGACA	CGACCGTGCT	TTAAAAACTT	GCTCTCCATT	420
	ATGTACTTCC	TTCCATCAGG	TTGGGGAAAA	AAAAATGGTG	GGGATGGTGA	GTAAACACAC	480
30	CAGTGGTTTC	ATCAGAGGG	AACTCACTAC	TCAGGAGGTG	ACGGTGACGT	GGTGCCGGTC	540
30	CCTGAAGTAC	GCGCACAAGC	TCCGGAGGTT	GCGGGAGCTT	CCGCTGCCGC	CTGGAGGGAA	600
	GCCGGAGCGA	CGGGGGTCAC	GGCGGCGGTC	AGAGGGTAAA	GGTCTTGCTC	CCAGCAGCCT	660
35	CCGCGGTGGA	TACGTCGCCA	TCTTGGATCC	GCGGGACAAG	AAAATTCATG	CGAGGGAGAC	720
	GTGGTGGGCG	GTCCTTCCTG	TGACACGACC	CTTGAGTGAC	AGTTCTATTT	GATTGCCTCC	780
40	GGTACTGTGA	GGAAAGGACA	CGACTCTATG	GTGAGGACTG	ATGGACATAC	ATTATCTGAG	840
10	AAAAGAAACT	ACCAGGTGAC	AAACAGCATG	TTTGGTGCTT	CAAGAAAGAA	GTTTGTAGAG	900
	GGGGTCGACA	GTGACTACCA	TGACGAAAAC	ATGTACTACA	GCCAGTCTTC	TATGTTTCCA	960
45	CATCGGTCAG	AAAAAGATAT	GCTGGCATCA	CCATCTACAT	CAGGTCAGCT	GTCTCAGTTT	1020
	GGGGCAAGTT	TATACGGGCA	. ACAAAGTGC	CTAGGCCTTC	CAATGAGGG	GATGAGCAAC	1080
50	AATACCCCTC	AGTTAAATCG	CAGCTTATCA	CAAGGCACTO	AGTTACCGAC	CCACGTCACG	1140
50	CCAACAACAG	GGGTACCAAC	AATGTCACTT	CACACGCCTC	CATCTCCAAC	CAGGGGTATT	1200
	TTGCCTATGA	ATCCTARGAA	TATGATGAA	CACTCCCAGG	TTGGTCAGG	G CATTGGAATT	1260
55	CCTAGCAGGA	CAAATAGCAT	GAGCAGTTC	A GGGTTAGGTA	A GCCCCAACA(AAGCTCGCCA	1320
	AGCATAATAT	GTATGCCAA	GCAGCAGCC	TCTCGACAG(CTTTACTG	T GAACAGTATG	1380
60	TCTGGATTTC	GAATGAACAC	GAATCAGGC	A TTTGGAATG	A ATAACTCCT	T ATCAAGTAAC	1440

	ATTTTTAATG	GAACAGACGG	AAGTGAAAAT	GTGACAGGAT	TGGACCTITC	AGATTTCCCA	1500
	GCATTAGCAG	ACCGAAACAG	GAGGGAAGGA	AGTGGTAACC	CAACTCCATT	AATAAACCCC	1560
5	TTGGCTGGAA	GAGCTCCTTA	TGTTGGAATG	GTAACAAAAC	CAGCAAATGA	ACAATCCCAG	1620
	GACTTCTCAA	TACACAATGA	AGATTTTCCA	GCATTACCAG	GCTCCAGCTA	TAÀAGATCCA	1680
10	ACATCAAGTA	ATGATGACAG	TAAATCTAAT	TTGAATACAT	CTGGCAAGAC	AACTTCAAGT	1740
10	ACAGATGGAC	CCAAATTCCC	TGGAGATAAA	AGTTCAACAA	CACAAAATAA	TAACCAGCAG	1800
	AAAAAAGGGA	TCCAGGTGTT	ACCTGATGGT	CGGGTTACTA	ACATTCCTCA	AGGGATGGTG	1860
15	ACGGACCAAT	TTGGAATGAT	TGGCCTGTTA	ACATTTATCA	GGGCAGCAGA	GACAGACCCA	1920
	GGAATGGTAC	ATCTTGCATT	AGGAAGTGAC	TTAACAACAT	TAGGCCTCAA	TCTGAACTCT	1980
20	CCTGAAAATC	TCTACCCCAA	ATTTGCGTCA	CCCTGGGCAT	CTTCACCTTG	TCGACCTCAA	2040
20	GACATAGACT	TCCATGTTCC	ATCTGAGTAC	TTAACGAACA	TTCACATTAG	GGATAAGCTG	2100
	GCTGCAATAA	AACTTGGCCG	ATATGGTGAA	GACCTTCTCT	TCTATCTCTA	TTACATGAAT	2160
25	GGAGGAGACG	TATTACAACT	TTTAGCTGCA	GTGGAGCTTT	TTAACCGTGA	TTGGAGATAC	2220
	CACAAAGAAG	AACGAGTATG	GATTACCAGG	GCACCAGGCA	TGGAGCCAAC	AATGAAAACC	2280
30	AATACCTATG	AGAGGGGAAC	ATATTACTTC	TTTGACTGTC	TTAACTGGAG	GAAAGTAGCT	2340
	AAGGAGTTCC	ATCTGGAATA	TGACAAATTA	GAAGAACGGC	CTCACCTGCC	ATCCACCTTC	2400
	AACTACAACC	CTGCTCAGCA	AGCCTTCTAA	АААААААА	ААААААААА	AAAAAGACTT	2460
35	CCCTTTTCTT	GGGGTATGGC	TGTCTCAGCA	CAATACTCAA	CATAACTGCA	GAACTGATGT	2520
	GGCTCAGGCA	CCCTGGTTTT	AATTCCTTGA	GGATCTGGCA	ATTGGCTTAC	GCAAAAGGTC	2580
40	ACCATTTGAG	GTCCTGCCTT	ACTAATTATG	TGCTGCCCAA	CAACTAAATT	TGTAATTTGT	2640
	TTTTCTCTAG	TTTGAGCAGG	GTCTGAATTT	TTTCATTTAT	TICCTTITIT	GCCAGCAGAC	2700
	AGACTTGAGT	CTGTAAAGAC	AAGCAAATAC	ACTGACAGAA	GITTACCATA	GTTTCTAAAA	2760
45	TGTAAAAAAG	AAAACCCCCA	AAAGACTCAA	GAAAATTAGA	. CCACAAATTT	TGCATTGTTC	2820
	ATTGTAGCAC	TATTGGTAAT	AAAATAACAA	ATGTTTGTGC	ATTTTTATGT	GAAGATCCTT	2880
50	CTCGTATTTC	ATTTGGAAAG	ATGAGCAAGA	GGTCTGCTTC	CTTCATTTTA	CTTCCCCTTC	2940
	TGTTTTTGAA	AGGCAGTTTC	GCCAAGCTTA	ATGCAAGAAT	ATCTGACTGT	TTAGAAGAAA	3000
	GATATTGCCA	CAATCTCTGG	ATGGTTTTCC	AGGGTTGTGT	TATTACTGAG	CTTCATCTTT	3060
55	CCAGAATGAG	CAAAACACTG	TCCAGTCTTT	GTTACGATTI	TGTAATAAAT	GTGTACATTT	3120
	TTTTTAAATT	TTTGGACATC	ACATGAATAA	AGGTATGTAI	GTACGAATGI	GTATATATTA	3180
60	TATATATGAC	ATCTATTITG	GAAAATGTTT	GCCCTGCTGT	ACCTCATTT	TAGGAGGTGT	3240

	GCATGGATGC	AATATATGAA	AATGGGACAT	TCTGGAACTG	CTGGTCAGGG	GACTTTGTCG	3300
	CCCTGTGCAC	TAAAAGGGCC	AGATTTTCAG	CAGCCAAGGA	CATCCATACC	CAAGTGAATG	3360
5	TGATGGGACT	TAAAAGAAGT	GAACTGAGAC	AATTCACTCT	GGCTGTTTGA	ACAGCAGCGT	3420
	TTCATAGGAA	GAGAAAAAA	GATCAATCTT	GTATTTTCTG	ACCACATAAA	GGĊTTCTTCT	3480
10	CTTTGTAATA	AAGTAGAAAA	GCTCTCCTCA	АААААААА	AAAAAAACTC	GAG	3533

(2) INFORMATION FOR SEQ ID NO: 25:

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- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1148 base pairs

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:

- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

ACCCACGCGT CCGCAAATTA TACTTCCTCA TTCATATTAT GTTGATACAA AAGACCTTGG 60 CAGCCATTTC TCCCAGCAGT TTTAAAGGAT GAACATTGGA TTTCATGCCA TCCCATAGAA 120 AACCTGTTTT AAAATTTTAG GGATCTTTAC TTGGTCATAC ATGAAAAGTA CACTGCTTAG 180 AAATTATAGA CTATTATGAT CTGTCCACAG TGCCCATTGT CACTTCTTTG TCTCATTTCT 240 300 TCCCTTTGTT CCTTAGTCAT CCAAATAAGC CTGAAAACCA TAAGAGATAT TACTTTATTG AATATGGTTG GCATTAAATT TAGCATTTCA TTATCTAACA AAATTAATAT AAATTCCAGG 360 420 ACATGGTAAA ATGTGTTTTA ATAACCCCCA GACCCAAATG AAAATTTCAA AGTCAATACC 480 AGCAGATTCA TGAAAGTAAA TTTAGTCCTA TAATTTTCAG CTTAATTATA AACAAAGGAA CAAATAAGTG GAAGGGCAGC TATTACCATT CGCTTAGTCA AAACATTCGG TTACTGCCCT 540 TTAATACACT CCTATCATCA GCACTTCCAC CATGTATTAC AAGTCTTGAC CCATCCCTGT 600 CGTAACTCCA GTAAAAGTTA CTGTTACTAG AAAATTTTTA TCAATTAACT GACAAATAGT 660 720 TTCTTTTTAA AGTAGTTTCT TCCATCTTTA TTCTGACTAG CTTCCAAAAT GTGTTCCCTT 780 TTTGAATCGA GGTTTTTTTG TTTTGTTTTG TTTTCTGAAA AAATCATACA ACTTTGTGCT TCTATTGCTT TTTTGTGTTT TGTTAAGCAT GTCCCTTGGC CCAAATGGAA GAGGAAATGT 840 900 TTAATTAATG CTTTTTAGTT TAAATAAATT GAATCATTTA TAATAATCAG TGTTAACAAT 960 TTAGTGACCC TTGGTAGGTT AAAGGTTGCA TTATTTATAC TTGAGATTTT TTTCCCCTAA 1020 CTATTCTGTT TTTTGTACTT TAAAACTATG GGGGAAATAT CACTGGTCTG TCAAGAAACA 1080 GCAGTAATTA TTACTGAGTT AAATTGAAAA GTCCAGTGGA CCAGGCATTT CTTATATAAA

TAAAATTGGT GGTACTAATG TGAAAAAAAA AAAAAAAAA AACTCGAGGG GGGCCCGGTA

	CCCTATTA	1148
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	(2) INFORMATION FOR SEQ ID NO: 26:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 717 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:	
	GGCACGAGCT AGCTGCCGCC ACCCGAACAG CCTGTCCTGG TGCCCCGGCT CCCTGCCCCG	60
20	CGCCCAGTCA TGACCCTGCG CCCCTCACTC CTCCCGCTCC ATCTGCTGCT GCTGCTGCTG	120
20	CTCAGTGCGG CGGTGTGCCG GGCTGAGGCT GGGCTCGAAA CCGAAAGTCC CGTCCGGACC	180
	CTCCAAGTGG AGACCCTGGT GGAGCCCCCA GAACCATGTG CCGAGCCCGC TGCTTTTGGA	240
25	GACACGCTTC ACATACACTA CACGGGAAGC TIGGTAGATG GACGTATTAT TGACACCTCC	300
	CTGACCAGAG ACCCTCTGGT TATAGAACTT GGCCAAAAGC AGGTGATTCC AGGTCTGGAG	360
30	CAGAGTCTTC TCGACATGTG TGTGGGAGAG AAGCGAAGGG CAATCATTCC TTCTCACTTG	420
30	GCCTATGGAA AACGGGGATT TCCACCATCT GTCCCAGCGG ATGCAGTGGT GCAGTATGAC	480
	GTGGAGCTGA TTGCACTAAT CCGAGCCAAC TACTGGCTAA AGCTGGTGAA GGGCATTTTG	540
35	CCTCTGGTAG GGATGGCCAT GGTGCCAGCC CTCCTGGGCC TCATTGGGTA TCACCTATAC	600
	AGAAAGGCCA ATAGACCCAA AGTCTCCAAA AAGAAGCTCA AGGAAGAGAA ACGAAACAAG	660
40	AGCAAAAGA AATAATAAAT AATAAATTT AAAAAAAAA AAAAAAA	717
45	(2) INFORMATION FOR SEQ ID NO: 27: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1099 base pairs (B) TYPE: nucleic acid	
50	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:	
	GGCACGAGCC GATGTGGACA TCATCCTGTC TATCCCCATG TTCCTGCGCC TGTACCTGAT	 60
55	CGCCCGAGTC ATGCTGCTGC ACAGCAAGCT CTTCACCGAT GCCTCGTCCC GCAGCATCGG	
	GGCCCTCAAC AAGATCAACT TCAACACCCG CTTTGTCATG AAGACGCTCA TGACCATCTG	120
60	·	180
00	CCCTGGCACT GTGCTGCTCG TGTTCAGCAT CTCTCTGTGG ATCATTGCTG CCTGGACCGT	240

	CCGTGTCTGT	GAAAGTCCTG	AATCACCAGC	CCAGCCTTCT	GGCTCATCAC	TTCCTGCTTG	300
5	GTACCATGAC	CAGCAGGACG	TAACTAGTAA	CTTTCTGGGT	GCCATGTGGC	TCATCTCCAT	360
	CACATTCCTT	TCCATTGGTT	ATGGGGACAT	GGTGCCCCAC	ACATACTGTG	GGAAAGGTGT	420
	CTGTCTCCTC	ACTGGCATCA	TGGGTGCAGG	CTGCACTGCC	CTTGTGGTGG	CCGTGGTGGC	480
10	CCGAAAGCTG	GAACTCACCA	AAGCGGAGAA	GCACGTTCAT	AACTTCATGA	TGGACACTCA	540
	GCTCACCAAG	CGGATCAAGA	ATGCTGCAGC	CAATGTCCTT	CGGGAAACAT	GGTTAATCTA	600
15	TAAACACACA	AAGCTGCTAA	AGAAGATTGA	CCATGCCAAA	GTGAGGAAAC	ACCAGAGGAA	660
13	GTTCCTCCCA	AGCTATCCAC	CAGTTTGAGG	AGCGTCCCAG	ATGGAACAGA	GGAAAGCTGA	720
	GTGACCAAGC	CAACACTCTG	GTGGACCTTT	CCAAGATGCA	GAATGTCATG	TATGACTTAA	780
20	TCACAGAACT	CAATGACCGG	AGCGAAGACC	TGGAGAAGCA	GATTGGCAGC	CTGGAGTCGA	840
	AGCTGGAGCA	TCTCACCGCC	AGCTTCAACT	CCCTGCCGCT	GCTCATCGCC	GACACCCTGC	900
25	GCCAGCAGCA	GCAGCAGCTC	CTGTCTGCCA	TCATCGAGGC	CCGGGGTGTC	AGCGTGGCAG	960
20	TGGGCACCAC	CCACACCCCA	ATCTCCGATA	GCCCCATTGG	GGTCAGCTCC	ACCTCCTTCC	1020
	CGACCCCGTN	CACAAGTTCA	AGCAGTTGCT	AAATAAATCT	CCCCACTCCA	GAAGCATTAA	1080
30	ААААААААА	АААААААА					1099

35 (2) INFORMATION FOR SEQ ID NO: 28:

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(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 941 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

45	AATTCGGCAG	AGAGCCAACC	GAGGGCGTTC	CTGTCGGGGC	TGCAGCGGCG	GGAGGGAGCC	60
	CAGTGGAGGC	GCCCTCCCGA	AGCGCCACTG	CCCATGCTGA	CCACCCAGCC	CTCCGGCTGC	120
50	TGATGTCATG	AGTAACACCA	CTGTGCCCAA	TGCCCCCCAG	GCCAACAGCG	ACTCCATGGT	180
	GGGCTATGTG	TTGGGGCCCT	TCTTCCTCAT	CACCCTGGTC	GGGGTGGTGG	TGGCTGTGGT	240
	AATGTATGTA	CAGAAGAAAA	AGCGGGTGGA	CCGGCTGCGC	CATCACCTGC	TCCCCATGTA	300
55	CAGCTATGAC	CCAGCTGAGG	AACTGCATGA	GGCTGAGCAG	GAGCTGCTCT	CTGACATGGG	360
	AGACCCCAAG	GTGGTACATG	GCTGGCAGAG	TGGCTACCAG	CACAAGCGGA	TGCCACTGCT	420
60	GGATGTCAAG	ACGTGACCTG	ACCCCCTTGC	CCCACCCTTC	AGAGCCTGGG	GTYCTGGACT	480

	GCCTGGGGCC	CTGCCATCTG	CTTCCCCTGC	TGTCACCTGG	STCCCCCTGC	TGGGTGCTGG	540
	GTCTCCATTT	CTCCCTCCAC	CCACCCTCAG	CAGCATCTGC	TTCCCATGCC	CTCACCATCA	600
5	CCTCACTGCC	CCCAGGCCTT	CTGCCCTTTG	TGGGTGTTGA	GCTCACCGCC	CACCCACAGG	660
	CACTCATGGG	AAGAGGCTTT	CCTTCTGGGA	TGGCGGCGGC	TGGTAGACAC	CTTTGCTTTC	720
10	TCTAGCCCTC	CTGGGCTGGG	CTTGGGCACA	AATCCCCAGG	CAGGCTTTGG	AGTTGTTTCC	780
	ATGGTGATGG	GGCCAGATGT	ATAGTATTCA	GTATATATTT	TGTAAATAAA	ATGTTTTGTG	840
	GCTAAAAAA	AAAAAAAA	ATCNAAGGGG	GGGCCGGTAC	CCAAATTCCC	CCTATANTGA	900
15	ATTCGTATTA	ACAATTCACT	TGGGGCCGTC	CTTTTAANAA	С		941

20 (2) INFORMATION FOR SEQ ID NO: 29:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 756 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

30	GGCACGAGGA AGCTGGAGCG GGCCGGCGGT GCAGTCACGG GGGAGCGAGG CCTGCTGGGC	60
	TTGGCAACGA GGGACTCGGC CTCGGAGGCG ACCCAGACCA CACAGACACT GGGTCAAGGA	120
35	GTAAGCAGAG GATAAACAAC TGGAAGGAGA GCAAGCACAA AGTCATCATG GCTTCAGCGT	180
	CTGCTCGTGG AAACCAAGAT AAAGATGCCC ATTTTCCACC ACCAAGCAAG CAGAGCCTGT	240
	TGTTTTGTCC AAAATCAAAA CTGCACATCC ACAGAGCAGA GATCTCAAAG ATTATGCGAG	300
40	AATGTCAGGA AGAAAGTTTC TGGAAGAGAG CTCTGCCTTT TTCTCTTGTA AGCATGCTTG	360
	TCACCCAGGG ACTAGTCTAC CAAGGTTATT TGGCAGCTAA TTCTAGATTT GGATCATTGC	420
45	CCAAAGTTGC ACTTGCTGGT CTCTTGGGAT TTGGCCTTGG AAAGGTATCA TACATAGGAG	480
.5	TATGCCAGAG TAAATTCCAT TTTTTTGAAG ATCAGCTCCG TGGGGCTGGT TTTGGTCCAC	540
	AGCATAACAG GCACTGCCTC CTTACCTGTG AGGAATGCAA AATAAAGCAT GGATTAAGTG	600
50	AGAAGGGAGA CTCTCAGCCT TCAGCTTCCT AAATTCTGTG TCTGTGACTT TCGAAGTTTT	660
	TTAAACCTCT GAATTTGTAC ACATTTAAAA TTTCAAGTGT ACTTTAAAAT AAAATACTTC	720
55	TAATGGAAAA AAAAAAAAAAAAAA ACTCGA	756

⁽²⁾ INFORMATION FOR SEQ ID NO: 30:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2100 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

10	NCCAGAGGCA	GAAAGTCCTG	CTTCTGGGGC	GTAACCTACA	GGATATCCTT	GGAACAGAAG	60
10	ATCTTATTGT	GGAAGTRACT	TCCAATGATG	CTGTGAGATT	TTATCCCTGG	ACCATTGATA	120
	ATAAATACTA	TTCAGCAGAC	ATCAATCTAT	GTGTGGTGCC	AAACAAATTT	CTTGTTACTG	180
15	CAGAGATTGC	AGAATCTGTC	CAAGCATTTG	TGGTTTACTT	TGACAGCACA	CAAAAATCGG	240
	GCCTTGATAG	TGTCTCCTCA	TGGCTTCCAC	TGGCAAAAGC	ATGGTTACCY	GAGGTGATGA	300
20	TCTTGGTCTG	CGATAGAGTG	TCTGAAGATG	GTATAAACCG	ACAAAAAGCT	CAAGAATGGT	360
20	GCATCCAAAC	ATGGCTTTGA	ATTGGTAGAA	CTTAGTCCAG	AGGAGTTGCC	TGAGGAGGAT	420
	GATGACTTCC	CAGAATCTAC	AGGAGTAAAG	CGAATTGTCC	AAGCCCTGAA	TGCCAATGTG	480
25	TGGTCCAATG	TAGTGATGAA	GAATGATAGG	AACCAAGGCT	TTAGCTTGCT	GCAACTCATT	540
	GACTGGAACA	AACCATAGCA	TTGGGTCAGC	AGATCCCTGT	CACCCAGAGC	AACCCCATTT	600
30	GCCAGCAGCA	GATAGTACTG	AATCCCTCTC	TGATCATCGG	GGTGGTGCAT	CTAACACAAC	660
30	AGATGCCCAG	GTTGATAGCA	TTGTGGATCC	CATGTTAGAT	CTGGATATTC	AAGAATTAGC	720
	CAGTCTTACC	ACTGGAGGAG	GAGATGTGGA	GAATTTTGAA	AGACTCTTTT	CAAAGTTAAA	780
35	GCAAATGAAA	GACAAGGCTG	CGACGCTTCC	TCATGAGCAA	AGAAAAGTGC	ATGCAGAAAA	840
	GGTGGCCAAA	GCATTCTGGA	TGGCAATCGG	GGGAGACAGA	GATGAAATTG	AAGGCCTTTC	900
40	ATCTGATGAA	GAGCACTGAA	TTATTCATAC	TAGGGTTTGA	CCAACAAAGA	TGCTAGCTGT	960
40	CTCTGAGATA	CCTCTCTACT	CAGCCCAGTC	ATATTTTGCC	AAAATTGCCC	TTATCATGTT	1020
	GGCTGCCTGA	CTTGTTTATA	GGGTCCCCTT	AATTTTAGTT	TTTAGTAGGA	GGTTAAGGAG	1080
45	AAATCTTTTT	TTTCCTCAGT	ATATTGTAAG	AGAGTGAGGA	ATACAGTGAT	AGTAATGAGT	1140
	GAGGATTTCT	TAAATRTACT	TTTTTTTTGT	TCTAGGAATG	AGGGTAGGAT	AAATCTCAGA	1200
50	GGTCTGTGTG	ATTTACTCAA	GTTGAAGACA	ACCTCCAGGC	CATTCCTGGT	CAACCTTTTA	1260
30	AGTAGCATTT	CCAGCATTCA	CACTTGATAC	TGCACATCAG	GAGTTGTGTC	ACCTTTCCTG	1320
	GGTGATTTGG	GTTTTCTCCA	TTCAAGGAGC	TTGTAGCTCT	GAAGCTATGA	TGCTTTTATT	1380
55	GGGAGGAAAG	GAGGCAGCTG	CAGAATTGAT	GTGAGCTATO	TGGGGCCGAA	GTCTCAGCCC	1440
	GCAGCTAAGT	CTCTACCTAA	. GAAAATGCCT	CTGGGCATTC	TTTTGAAGT#	TAGTGTCTGA	1500
60	GCTCATGCTA	. GAAAGAATCA	AAAAGCCAGT	GTGGATTTT	C AGACTGTAAT	· AAATGAGGCA	1560

	AAGGATTTCT	ATTCCAGTGG	GAAGRAAACC	TCTCTACTGA	GTTGTGGGG	ATATGTTGTA	1620
	TGTTAGAGAG	AACCTTAAGG	AGTCCTTGTA	TGGGCCATGG	AGACAGTATG	TGATAACATA	1680
5	CCGTGATTTT	CATGAAGAAA	TTCTTCTGTC	TTAGAGTTCT	CCCCTGCTGC	TTGAGATGCC	1740
	AGAGCTGTGT	TGTTGCACAC	CTGCAAAACA	AGGCACATTT	CCCCCTTTCT	CTTTAAAGCC	1800
10	AAAGAGAGAT	CACTGCCAAA	GTGGGAGCAC	TAAGGGGTGG	GTGGGGAAGT	GAAATGTTAG	1860
10	GCGATGAATT	CCTGAGCACC	TIGTTTTTCT	TCCAAGGTTC	GTAGCTCCTC	TCTGCCCTTC	1920
	CAAGCCTGTA	ACCTCGGAGG	ACTATCTTTT	GTTCTTTATC	CTTTGTCTTG	TTTGAGTGGG	1980
15	TCAGCCCCAG	AGGAACTGAT	AAGCAAATGG	CAAGTTTTTA	AAGGAAGAGT	GGAAAGTACT	2040
	GCAAATAAAA	ATCCTTATTT	GTTTTTGTAG	АААААААА	ААААААААА	AAAAAAAAG	2100

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(2) INFORMATION FOR SEQ ID NO: 31:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1448 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:

ааааааааа	AAAGCCCACC	TGAAAGCCTG	TCTCTTTCCA	CTTTGTTGGC	CCTTCCAGTG	60
GGATTATCGA	GCATGTTGTT	TTTTCATAGT	GCCTTTTTCC	TTATTTCAAG	GGTTGCTTCT	120
GAGTGGTGTT	TTTTTTTTT	TTAATTTGTT	TTGTTTTAAA	ATAAGTTAAA	GACAGTCCAG	180
AGCTTTTCAG	CCAATTTGTC	TCCTACTCTG	TGTAAATATT	TTTCCCTCCG	GGCAGGGGAG	240
CCAGGGTAGA	GCAAAGGAGA	CAAGCAGGAG	TGGAAGGTGA	GGCGTTCTCC	TGCTTGTACT	300
AAGCCAGGAG	STTTAAGCTC	CAGCTTTAAG	GGTTGTGAGC	CCCTTGGGGT	TCAGGGAACT	360
GCTTGCCCAG	GGTGCAGTGT	GAGTGTGATG	GCCACCGG	GCAAGAGGGA	AGGTGACCGC	420
CCAGCTCTCC	CACATCCCAC	TGGATCTGGC	TTACAGGGG	GTCGGAAGCC	TGTCCTCACC	480
CTCTCGGGG	TTGTGGCCCC	CCCCCCTCC	CTATATGCAC	CCCTGGAACC	AGCAAGTCCC	540
AGACAAGGAG	AGCGGAGGAG	GAAGTCATGG	GAACGCAGCC	TCCAGTTGTA	GCAGGTTTCA	600
СТАТТССТАТ	GCTGGGGTAC	ACAGTGAGAG	TACTCACTTT	TCACTTGTCT	TGCTCTTAGA	660
TTGGGCCATG	GCTTTCATCC	TGTGTCCCCT	GACCTGTCCA	GGTGAGTGTG	AGGGCAGCAC	720
TGGGAAGCTG	GAGTGCTGCT	TGTGCCTCCC	TTCCCAGTGG	GCTGTGTTGA	CTGCTGCTCC	780
CCACCCCTAC	CGATGGTCCC	AGGAAGCAGG	GAGAGTTGGG	GAAGGCAAGA	TTGGAAAGAC	840
AGGAAGACCA	AGGCCTCGGC	AGAACTCTCT	GICTTCTCTC	CACTTCTGGT	CCCCTGTGGT	900

	GATGTGCCTG TAATCTTTTT CTCCACCCAA ACCCCTTCCC ACGACAAAA CAAGACTGCC	960
5	TCCCTCTCTT CCGGGAGCTG GTGACAGCCT TGGGCCTTTC AGTCCCAAAG CGGCCGATGG	1020
3	GAGTCTCCCT CCGACTCCAG ATATGAACAG GGCCCAGGCC TGGAGCGTTT GCTGTGCCAG	1080
	GAGGCGGCAG CTCTTCTGGG CAGAGCCTGT CCCCGCCTTC CCTCACTCTT CCTCATCCTG	1140
0	CTTCTCTTTT CCTCGCAGAT GATAAAAGGA ATCTGGCATT CTACACCTGG ACCATTTGAT	1200
	TGTTTTATTT TGGAATTGGT GTATATCATG AAGCCTTGCT GAACTAAGTT TTGTGTGTAT	1260
15	ATATTTAAAA AAAAAATCAG TGTTTAAATA AAGACCTATG TACTTAATCC TTTAACTCTG	1320
15	CGGATAGCAT TTGGTAGGTA GTGATTAACT GTGAATAATA AATACACAAT GAATTCTTMA	1380
	AAAAAAAAA AAAAAAAAA AAAAAAAAA AAACCCCGGG GGGGCCCCG GGCCCCAATT	1440
20	ССССССАА	1448
25	(2) TYPODYARTON FOR CEO TO NO. 22.	
2.5	(2) INFORMATION FOR SEQ ID NO: 32:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 456 base pairs	
30	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:	
35	GGCACAGCAA ACTTGACGCC ATGAAGATCC CGGTCCTTCC TGCCGTGGTG CTCCTCTCCC	60
	TCCTGGTGCT CCACTCTGCC CAGGGAGCCA CCCTGGGTGG TCCTGAGGAA GAAAGCACCA	120
40	TTGAGAATTA TGCGTCACGA CCCGAGGCCT TTAACACCCC GTTCCTGAAC ATCGACAAAT	180
	TGCGATCTGC GTTTAAGGCT GATGAGTTCC TGAACTGGCA CGCCCTCTTT GAGTCTATCA	240
	AAAGGAAACT TCCTTTCCTC AACTGGGATG CCTTTCCTAA GCTGAAAGGA CTGAGGAGCG	300
45	CAACTCCTGA TGCCCAGTGA CCATGACCTC CACTGGAAGA GGGGGCTAGC GTGAGCGCTG	360
	ATTCTCAACC TACCATAACT CTTTCCTGCC TCAGGAACTC CAATAAAACA TTTTCCATCC	420
50	AAAAAAAAA AAAAAAAAC CCCNGGGGGG GCCCGG	456
	•	
•	(2) INFORMATION FOR SEQ ID NO: 33:	
55		
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1326 base pairs	
	(B) TYPE: nucleic acid	

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

5	GGCACGAGTG CAGGCCCAGA GAGGACTCAT TGAAAGGACT GAAAGGGGAG GTGGCGTTTT	60
J	CTTCCTACCC AAACTTACCC CTGTGAGCTG GACAGCTTGG TAGCACCTGC CTGGACTTAG	120
	ATGGTGGTAG CCAAGAAGAC TGACATTTTA GGGAACAGGA CGGGGAGGAG AAGGCTCTGG	180
10	CACACACA TGTGTCCATA TGTCCTGCAA TGGTCTGGGG ACTATTGCTA GGCTAGGAGC	240
	CCTAAGTGTC TTCTTCCTCA TGTCTMTTCT CCCCTGTSTC ATGGGCCCTA AGRTCTCTTT	300
15	CACTGGGCCT GCCTCAATGA ACGTGCTGCC CAGCTACCCC GAAACACGGC ANCTGCCGGC	360
15	TATCAATGCC CCAGCTGCAA TGGCCCATCT TCCCCCAACC AACCTGGCTG GGCCCGTGGG	420
	CTCCGCACTG AGARARAAAS TTGGCACART CAACTGGGCC CGGGCAGGAC TGGGCCYCCC	480
20	TCTGATCGAT GAAGKTGGTG ARCCCAGAGC CCGAGCCCCT CAACACGTCT GACTTCTCTG	540
	ACTGGTCTAG TTTTAATGCC AGCAGTACCC CTGGACCAGA GGAGGTAGAC AGCGCCTCTG	600
25	CTGCCCCAGC CTTCTACAGC CGAGCCCCCC GGCCCCAGC TTCCCCAGGC CGGCCCGAGC	660
	AGCACACAGT GATCCACATG GGCAATCCTG AGCCCTTGAC TCACGCCCCT AGGAAGGTGT	720
	ATGATACGCG GGATGATGAC CGGACACCAG GCCTCCATGG AGACTGTGAC GATGACAAGT	780
30	ACCGACGTCG GCCGGCCTTG GGTTGGCTGG CCCGGCTGCT AAGGAGCCGG GCTGGGTCTC	840
	GGAAGCGRCC GCTGACCCTG CTCCAGCGGG CGGGGCTGCT GCTACTCTTG GGACTGCTGG	900
35	GCTTCCTGGC CCTCCTTGCC CTCATGTCTC GCCTAGGCCG GGCCGCAGCT GACAGCGATC	960
	CCAACCTGGA CCCACTCATG AACCCTCACA TCCGCGTGGG CCCCTCCTGA GCCCCCTTGC	1020
	TTGTGGCTAG GCCAGCCTAG GATGTGGGTT CTGTGGAGGA GAGGCGGGGT AATGGGGAGG	1080
40	CTGAGGGCAC CTCTTCACTG CCCCTCTCCC TCAAGCCTAA GACACTAAGA CCCCAGACCC	1140
	AAAGCCAAGT CCACCAGAGT GGCTGCAGGC CAGGCCTGGA GTCCCCGTGG GTCAAGCATT	1200
45	TGTCTTGACT TGCTTTCCTC CCGGGTYTCC AGCCTCCGAC CCCTCGCCCC ATGAAGGAGC	1260
	TGGCAGGTGG AAATAAACAA CAACTTTATT AAAAAAAAAA	1320
	AAANAA	1326

50

(2) INFORMATION FOR SEQ ID NO: 34:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 710 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:	
	GCGAAAGAGA AAAAGGCTGG AGCTCCCGCC CCCGGGGCTG TCAGATGGCT TGGGTTTCTG	60
5	CGACGCGATT GGCTCGCGGA GGGCAGAAAT TACTCAGCAA ACATGACTAT TATTAGCTGC	120
	TTAGCAACAG CTCACCAAAG TAGAGAGACC ACCCAGGTAG GCAACCCAGT GTGTGCATCC	180
10	TCGGCTTCGG GGCAGCCTCT GAGAGCGCCA ACCTTCTCGC ATGCAATACT TCCATTAAGG	240
10	AATGCTCCCC CTCCTTTCTC TCTTATTCCT TTTCTTTTCA ACAGTGTCTT CTTTTTGTGG	300
	GATGCCTTTG CGCGCACACA CGCGCGCGCA SGCACACACA CGAACATTTG CCTCGCGGTA	360
15	GACACGGGG GAAATGTWAT ATTTTTTAA GCGCTTAAAC AATTTCTGAA ATTCCTCAAA	420
	GAAAAGCCTT TCAGARGCAC CTTGGCCTCA AGCTGCAACA AATACTGGGA RGTCCGGCTC	480
20	GCATTCCCAG GCCTGCACCA ATAATGACAG CGTGCTGGAT ARTGCGCCAG TGTGTGCCAG	540
20	ATTTTTTTT CCTCTTCTCT TTTCTTTAT AACTAAAGGG AAGACTTAGG CTCTTGCAGG	600
	GAACAACGCC TCGCATTAAG ATAAACAGAA TGGAAAGTTA AAGAGGAAAG CAAGGACGTT	660
25	GGGAAAAGCC ATCTTTCTTA AAATCCGTCT GCCCCCCAGC CGCTTTCTCC	710
30	(2) THE PROPERTY AND ADD ADD ADD ADD ADD ADD ADD ADD ADD	
50	(2) INFORMATION FOR SEQ ID NO: 35:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1188 base pairs	
35	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:	
40	GATGGCTTTT ATATCTATTA TCGACCCACA GACAGTGACA ATGATAGTGA CTACAAGAAG	60
	GATATGGTGG AAGGGGACAA GTACTGGCAC TCCATCAGCC ACCTGCAGCC AGAGACCTCC	120
45	TACGACATTA AGATGCAGTG CTTCAATGAA GGAGGGGAGA GCGAGTTCAG CAACGTGATG	180
	ATCTGTGAGA CCAAAGCTCG GAAGTCTTCT GGCCAGCCTG GTCGACTGCC ACCCCCAACT	240
	CTGGCCCCAC CACAGCCGCC CCTTCCTGAA ACCATAGAGC GGCCGGTGGG CACTGGGGCC	300
50	ATGGTGGCTC GCTCCAGCGA CCTGCCCTAT CTGATTGTCG GGGTCGTCCT GGGCTCCATC	360
	GTTCTCATCA TCGTCACCTT CATCCCCTTC TGCTTGTGGA GGGCCTGGTC TAAGCAAAAA	420
55	CATACAACAG ACCTGGGTTT TCCTCGAAGT GCCCTTCCAC CCTCCTGCCC GTATACTATG	480
	GTGCCATTGG GAGGACTCCC AGGCCACCAG GCAGTGGACA GCCCTACCTC AGTGGCATCA	540
	GTGGACGGC CTGTGCTAAT GGGATCCACA TGAATAGGGG CTGCCCCTCG GCTGCAGTGG	600
60	GCTACCCGGG CATGAAGCCC CAGCAGCACT GCCCAGGCGA GCTTCAGCAG CAGAGTGACA	660

GCTACCCGGG CATGAAGCCC CAGCAGCACT GCCCAGGCGA GCTTCAGCAG CAGAGTGACA

TCACGAGGGG TCCCAAGTCT AGCCCGACA AGGGCTCTTT CTTATAC ACTCCACTCA CCAGCTGCTG CAGCCCCATC ACGACTGCTG CCAACGC CTGSTGTGGG CCAGTCAGGG GTGAGGAGAG CCCCCGACAG TCCTGTC TGGACAGTCC TGACTCCTGC CAAGTGAGTG GAGGAGACTG GTGTCCC GGGCCTACGT AGGACAGGAA CCTGGAATGC AGCTCTCCCC GGGGCCA TTGTTTTTTT TTTAAAAAAA AAAAAAAAA AWCYCGGGGG GGGCCCC (2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDMESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG CCAGGAATCA CTCTGCTAGC AGAATGGCG CCCCATCCCT TACTATC AAGTGCAGCC CAGAAGGAC CAGGCCTTTG ATGCACATTG GGTGGGT ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGGTC ATTGACACTG GGCCACCCCC ACAACTTCTC CTCAAGGAG GTTNTCC GCAAAATGCC GAGCACTACT GCATAAGAAT TCTTCCTGCT TCTGGTC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC GCAAAATGCC GAGCACTACT GCATAAGAAT TCTTCCTGCT GTACTGC TTTGACACAG CCACCACCCC ACAACTTCTC CTCAAGGGAG GTTNTCC GCAAAATGCC GAGCACTACT GCATAAGGATG CCAGGACCAC CGAAGGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTTT GCCAGGACCA TTCCATCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 50 CTCCAGCACA TTCCATCAGG GCCTGTTTGAG CACCTTCTGA AAAGCAC 51 CTCCAGCACA TTCCATCAGG GCCTGTTTGAG CACCTTCTGA AAAGCAC 52 CTCCAGCACA TTCCATCAGG GCCTGTTTGAG CACCTTCTGA AAAGCAC 53 CTCCAGCACA TTCCATCAGG GCCTGTTTGAG CACCTTCTGA AAAGCAC 54 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGGT TTTAATAATA ATAAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTTT GCCAGATGCA CATCTTCAGG GCCTGTTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGGT TTTCATCAGCACA TTCCATCAGA GTCAGGAAAAC CTGCGGTGAG TCCCAGGT TTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGGT TTCCAGCACAC TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGGT TTCCAGCACAC TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCC	CCAA AGTCACCAGA	720
ACTCCACTCA CCAGCTGCTG CAGCCCCATC ACGACTGCTG CCAACACC CTGSTGTGGG CCAGTCAGGG GTGAGGAGAG CCCCCGACAG TCCTGTG CTGSTGTGGG CCAGTCAGGG GTGAGGAGAG CCCCCGACAG TCCTGTG GGGACCCTCC ATTTCACTCA GGGCCCCCAT GCTGCTTGGG CCTTGTGG TGGACAGTCC TGACTCCTGC CAAGTGAGTG GAGGAGACTG GTGTCCC GGGCCTACGT AGGACAGGAA CCTGGAATGC AGCTCTCCCC GGGGCCA TTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGCCC CTTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGCCC CTTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGCCCC CTTGTTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGCCCC CTTGTTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGCCCC CTTGTTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGCCCCCCCCCCCCCCCCCCCCCCCCCCC	CACA CTGCCCGACG	780
10 GGGACCTCC ATTTCACTCA GGGCCCCAT GCTGCTTGGG CCTTGTG TGGACAGTCC TGACTCCTGC CAAGTGAGTG GAGGAGACTG GTGTCCC GGGCCTACGT AGGACAGGAA CCTGGAATGC AGCTCTCCCC GGGGCCA CTTTTTGAAAC ACCACCTCT ACAATTTAGG CAGAAGCTGA TATCCCA TTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGCCC 20 (2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG 25 GATGTGACAT CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATC AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCACC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAACCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCCGTGAG TCCCAGG	CCAG GAGCAGCCTG	840
TGGACAGTCC TGACTCCTGC CAAGTGAGTG GAGGAGACTG GTGTCCC GGGCCTACGT AGGACAGGAA CCTGGAATGC AGCTCTCCCC GGGGCCA CTTTTGAAAAC ACCACCTCTC ACAATTTAGG CAGAAGCTGA TATCCCA TTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGGCCC (2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS:	CCTG GAAGCAGTGT	900
GGGCCTACGT AGGACAGGAA CCTGGAATGC AGCTCTCCCC GGGGCCA CTTTTGAAAC ACCACCTCTC ACAATTTAGG CAGAAGCTGA TATCCCA TTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGGCCC (2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS:	GCCA GTTGAAGAGG	960
CTTTTGAAAC ACCACCTCTC ACAATTTAGG CAGAAGCTGA TATCCCA TTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGGCCC (2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGACCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG 35 GATGTGACAT CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATC AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGG	CCAG CACCCCGTAG	1020
TTGTTTTTT TTTAAAAAAA AAAAAAAAAA AWCYCGGGGG GGGGCCC 20 (2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG 35 GATGTGACAT CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGGG CCCCATCCCT TACTATC AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAAGGAA AAATCTCAGC CTGCAGGACT CTGGTTT GCCAGGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGG	ACTG GTGCGTGTGT	1080
(2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG 35 GATGTGACAT CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGCG CCCCATCCCT TACTATC AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT AGTTGAAATG GGAGCATGCT GGAGTCGGC TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCACG TCCAGTATTC CATCCTTCCT TCTGTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC 50 GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGGACCA CATCCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGG	AGAA AGACTATATA	1140
(2) INFORMATION FOR SEQ ID NO: 36: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG CCAGGAATCA CTCTGCTAGC AGAATGGGC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATG AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGGTC ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGG	cc	1188
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (Xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATG CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATG AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCCGGTGAG TCCCAGG		
(A) LENGTH: 956 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG CCAGGAATCA CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATG AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGGTC ATTGACCCCT GGCCACCACG TCCAGTATTC CATCCTTCCT TCTGTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGACT CTGGTTT GCCAGGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGG		
(B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG CCAGGAATCA CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATG AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGGTC ATTGACCCCT GGCCACCACG TCCAGTATTC CATCCTTCCT TCTGTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAAATGTCC GAGCACTACT GCATAGGATG CCAGAGACT CTGGTTT GCCAGGACCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGG		
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGG GGCAGAGCAG TGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGGG CCCCATCCCT TACTATG AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC ATTGACCCCT GGCCACCACG TCCAGTATTC CATCCTTCCT TCTGTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCCGTGAG TCCCAGA		
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36: GGCAGAGCAG TGAAAAATGCA TCCTAAAAAAT TCAATGTTTA TACCAGG GGCAGAGCAG TGAAAAATGCA TCCTAAAAAAT TCAATGTTTA TACCAGG GATGTGACAT CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATC AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA		
GATGTGACAT CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTC CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATO AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCACG TCCAGTATTC CATCCTTCCT TCTGTCC TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA		
AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTM GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	GCTC ATGACACTAA	60
AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGT AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	CCCA GATTGCATCT	120
AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	GCTG CTCCTCCTCA	180
AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTC ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCC 45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	TCTC CCACTACTTT	240
45 TCCCCACYAG GCCCACCCCC ACAACTTCTC CTCAAGGGAG GTTNTCC CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	GAGA AGGACATCCC	300
CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGC GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	CCAG CCTATCGCCC	360
GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGAT TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAG 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	CCGC AGCTGGAGGG	420
TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAC 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	GGCC TTAACTGCCT	480
TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTT. GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAG 55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	TAAA CACAGCCAAG	540
55 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGA	TTGA CCCACCATCG	600
	GGGC TCGTAATAGA	660
1000010001 0010010M01 M110010M01 T110T1	AGAA TCTAGGGTGC	720
AGGGCAGGGA GCAGGAGTCA TAAGGAGTGA TAACCTAAAC TGTGTGT	TAGT CAGCGGGGAG	780
GGTCTTATGT TATCAGGTGA AATGAGAGCC AGTAAGTTAG TTGATCO	CTGT CACAGATATA	840

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1320

	ACCCTGATAA CACCCCATAG ATACGCGACA CGTGTGTCCT GCCCCTGCTT TCCCCATCCA	900
	ACATGGTTCT TCTGTTCCAC AGACATTAAA GGGGCTTTCT GCAATTACTT AAAAAA	956
5		
	·	
	(2) INFORMATION FOR SEQ ID NO: 37:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1603 base pairs(B) TYPE: nucleic acid	
1.5	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:	
	TCGACCCACG CGTCCGCTCT GCCAGGAATC TGGTCTTTCT GTAGACCCAA GTCAGAAAGA	60
20	ACCATTTGTG GAGTTAAATC GAATATTAGA RGCATTAAAR GTCAGAGTTC TGAGACCTGC	120
	TCTGGAATGG GCAGTTTCAA ACCGAGAGAT GCTTATAGCC CAAAACAGCT CCTTGGAATT	. 180
25	TAAACTACAC AGACTGTATT TTATTAGCTT RTTAATGGGT GGAACACAAA TCAGCGAGAR	240
23	GCATTACAAT ATGCTAAAAA TTTTCAGCCA TTTGCCCTAA ATCATCAAAA AGACATTCAG	300
	GTTTTGATGG GAAGCCTTGT GTACCTGAGA CAAGGGATTG AGAACTCACC ATATGTTCAC	360
30	CTACTIGATG CAAACCAGTG GGCTGATATC TGTGACATCT TTACACGGGA TGCTTGTGCC	420
	CTCCTGGGGC TCTCCGTGGA GTCCCCTCTC AGTGTCAGTT TCTCAGCAGG TTGTGTGGCG	480
35	CTGCCAGCTT TAATTAACAT CAAAGCCGTG ATTGAACAGA GGCAGTGTAC TGGAGTTTGG	540
<i>)</i>	AACCAGAAAG ATGAATTACC TATTGAAGTG GACCTTGGTA AAAAGTGCTG GTATCACTCT	600
	ATATTTGCCT GCCCCATTCT TCGTCAGCAA ACAACAGATA ACAATCCACC CATGAAATTG	660
40	GTCTGTGGTC ATATTATATC AAGAGATGCC CTGAATAAAA TGTTTAATGG TAGCAAATTA	720
	AAATGTCCCT ACTGTCCAAT GGAACAAAGT CCAGGAGATG CCAAACAGAT ATTTTTCTGA	780
45	AGAGATAACT TTAGTTTGCA ATTTGTAAGT GAAACTGAAT CGTGGGTGCA TTTCAGAAGA	840
43	GAACGTTCCA TATAATGCAG CTAACCAAGG ACTCCTGTGT TTCTATAAGC TAATGCTCCA	900
	GAAACTITGC CAACCTGTTA GTGTACACAC ACTGAGGGGA GTGCTCCCGG TGAATATTAT	960
50	CATAGGGCTT TATTATATTC TTGGTCTTCA TTTCTGATCA AGTAAATACA CCAGCAGTTG	1020
	TCATTCAATG CAGGITTITG TACTTAATTA TATGGTGATT TTTTTACTIT TTAAGAGCAG	1080
<i>e</i>	AAACGGAAAT TGACCTCCCC GCCATGTGTT TAATATTCCT CCTGCTTTTA CTTTTGTCAT	1140
55	TTTCTTGATA ATCGTAAGCC TTGAGAGTGT TTGTGAAAAA GTTTTATTTC CTGTTATGTA	1200
	TACATAATTA AATGAAAATT CTTCAGAAAA AGTTTGATAA ATTGAATTGT GGTTATGAAA	1260

CTAATTTGCA TITITATTTG CTTAAGAAAG AAAGCTGTGA TAGATTCCAG ATATGCTTTT

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	TGATGTTTTC	CTCTGCTCCA	GCTCCAAGAA	GTCAGCACAC	CTGCATTTTA	GCTCTGCATG	1380
5	CAGCCCCAGC	AGGCTGCGTG	TTTAAGAATT	TCATTGTTTA	ACTGGCTGGT	GTGAGAAGTC	1440
	TTCCGTTAGC	ATAGAGTGGA	AGGAGTACTA	TTGTTTGGTT	GGGTTTTTGT	TTGTTTGTTT	1500
	TTTGTTTTTG	CTTTTATTGC	CAAGAGGTGC	TTGTTTTAAA	AGTATGTTTA	ATAAAATGAA	1560
10	ATTCTAAAGT	TAARAAGTGT	TCTTAAAGTT	GATATTTAAC	TCT		1603

15 (2) INFORMATION FOR SEQ ID NO: 38:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1089 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

25	GGCACGAGCT	ACCTTTCTGC	CIGCTIIGCT	GGCTGCAACA	GCACGAATCT	CACGGGCTGT	60
	GCGTGCCTCA	CCACCGTCCC	TGCTGAGAAC	GCAACCGTGG	TTCCTGGAAA	ATGCCCCAGT	120
30	CCTGGGTGCC	AAGAGGCCTT	CCTCACTTTC	CTCTGTGTGA	TGTGTATCTG	CAGCCTGATC	180
50	GGTGCCATGG	CAAGACACCC	TCAGTCATCA	TCCTCATCAG	GACAGTCAGC	CCTGAACTCA	240
	AGTCTTACGC	TTTGGGAGTT	CTTTTTCTCC	TCCTTCGTTT	GTTGGGCTTC	ATCCCTCCAC	300
35	CCCTCATCTT	CGGGGCTGGC	ATCGACTCCA	CCTGCCTGTT	CTGGAGCACG	TTCTGTGGGG	360
	AGCAAGGCGC	CTGCGTCCTC	TACGACAATG	TGGTCTACCG	ATACCTGTAT	GTCAGCATCG	420
40	CCATCGCGCT	CAAATCCTTC	GCCTTCATCC	TGTACACCAC	CACGTGGCAG	TGCTGAGGAA	480
40	AAACTATAAA	CGCTACATCA	AAAACCACGA	GGCGGGCTG	AGCACCAGTG	AGTTCTTTGC	540
	CTCTACTCTG	ACCCTAGACA	ACCTGGGGAG	GGACCCTGTG	CCCGCAAACC	AGACACATAG	600
45	GACAAAGTTT	ATCTATAACC	TGGAAGACCA	TGAGTGGTGT	GAAAACATGG	AGTCCGTTTT	660
	ATAGTGACTA	AAGGAGGCT	GAACTCTGTA	TTAGTAATCC	AAGGGTCATT	TTTTTCTTAA	720
50	AAAAAGAAAA	AAAGGTTCCA	AAAAAAACCA	AAACTCAGTA	CACACACACA	GGCACAGATG	780
50	CACACACACA	CAGACAGACA	CACCGACTTT	GTCCTTTTC	TCAGCATCAG	AGCCAGACAG	840
	GATTCAGAAT	AAGGAGAGAA	TGACATCGTG	CGGCAGGGTC	CTGGAGGCCA	CTCGCGCGGC	900
55	TGGGCCACAG	AGTCTACTTT	GAAGGCACCI	CATGGTTTTC	AGGATGCTGA	CAGCTGCAAG	960
	CAACAGGCAC	TGCCAAATTC	AGGGAACAGT	GGTGGCCAGC	TTGGAGGATG	GACATTTCTG	1020
60	GATACACATA	CACATACAAA	ACAGAAAACA	AAAATTTTTT A	GAAGTTTCCI	' ААААТАААА	1080

	AAAAAAA	1089
5	(2) INFORMATION FOR SEQ ID NO: 39:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 629 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:	
13	AGCTCAGTTC CCTTAGAAAT GAAATTTTAA ATGACACTAC CAGGTAAGCC ACTGAGACCA	60
	GTGGAGGTGA TAGCTAAGAA CATAAGGAAT TAAGAATTTT TAATGGAGAA AGGAGGTAAT	120
20	GAATACCAGT TACATCCTAA GACTCACTGT AGTGGTGAGT GTTGTAATTT ATCTCGCTAT	180
	CCATCCTCTT TTAAGTTTTT CCTTAGAAAG TCCTCTATTG GTACCTTGGA GGGACTGCTG	240
25	TCAAAATATA TGGAAAAGTG GGTCTGTGTG GTACAAGAGG TGGACTTTGC CACACATGGA	300
	AGTTTGCTGC CAAGATCTTC ACTAATGAAA GAAATCACCA GTGAGCTGCA CAGATTAGCC	360
	AAATACTGAG CTCATTAGAA CTACTAAGGC CTGGACATTT CTGCCTAATC CAGGACTCCT	420
30	GTAATTATCA GTCTTTGCTT TGGAGCTTCC CATTGTGTAG CTGARAATTT GTCATATCTG	480
	CATTATAATC TAAGGCTCCA CATACTTAAT CCTGCTTCTC CCCCTTTTTC TTTCCCTTTC	540
35	CCAGCGGTCA GCTCTGCTGC ATAGTCTGAA GACTTTCCCT GCCCAATCCT GATAAAATTC	600
	TTGCACTCGT AACCCCATCT CAGTGTCTG	629
40	(2) INFORMATION FOR SEQ ID NO: 40:	
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1964 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:	
50	AAGAAGACAT GGAAATTGCT GAAGGATGTT TCAGGCATAT TAAGAAAATC TTTACGCAGC	60
	TTGAGGAATT CAGAGCCTCT GAATTGCTTC GAAGTGGACT GGACAGATCT AAATACCTTT	120
55	TAGTGAAAGA AGCCAAAATT ATTGCTATGA CCTGTACTCA TGCTGCCTTA AAACGACATG	180
	ACTIGGICAA GCTAGGITIC AAGTAIGACA ACATITIGAT GGAAGAGGCT GCTCAGATIC	240
60	TGGAGATAGA AACTTTTATC CCTCTTCTTC TACAGAATCC TCAGGATGGA TTTAGCCGAC	300

	TAAAACGATG GATTATGATT GGCGATCATC ACCAGTTACC TCCAGTTATT AANGAACATG	360
	GCCTTTCAAA AGTACTCAAA CATGGAGCAG TCTCTCTTCA CTCGCTTTGT TCGCGTTGGA	420
5	GTTCCGACTG TTGACCTTGA TGCTCAAGGG AGAGCCAGAG CAAGCTTGTG CAMCTNCTAC	480
	AACTGGCGAT ACAAGAATCT AGGAAACTTA CCCCATGTGC AGCTCTTGCC AGAGTTTAGT	540
10	ACAGCAAATG CTGGCTTACT GTATGACTTC CAGCTCATTA ATGTTGAAGA TTTTCAAGGA	600
10	GTGGGAGAAT CTGAACCTAA TCCTTACTTC TATCAGAATC TTGGAGAGGC AGAATATGTA	660
	GTAGCACTTT TTATGTACAT GTGTTTACTT GGTTACCCTG CTGACAAAAT CAGTATTCTA	720
15	ACAACATATA ATGGCCAAAA GCATCTTATT CGCGACATCA TCAATAGACG ATGTGGAAAC	780
	AATCCATTGA TTGGAAGACC AAACAAGGTG ACAACTGTTG ATAGATTTCA AGGTCAACAG	840
20	AATGACTATA TICTTCTTTC TCTGGTACGA ACCAGGGCAG TGGGCCATCT GAGGGATGTC	900
20	CGTCGCTTGG TAGTGGCCAT GTCTAGAGCC AGACTTGGAC TTTATATCTT CGCCAGAGTA	960
	TCCCTCTTCC AAAACTGTTT TGAACTGACT CCAGCTTTCA GTCAGCTCAC AGCTCGCCCC	1020
25	CTTCATTTGC ATATAATTCC AACAGAACCT TTCCCAACTA CTAGAAAGAA TGGAGAGAGA	1080
	CCATCTCATG AAGTACAAAT AATAAAAAAT ATGCCCCAGA TGGCAAACTT TGTATACAAC	1140
30	ATGTACATGC ATTTGATACA GACTACACAT CATTATCATC AGACTTTATT ACAACTACCA	1200
50	CCTGCTATGG TAGAAGAGGG TGAGGAAGTT CAAAATCAAG AAACAGAATT GGAAACAGAA	1260
	GAAGAGGCCA TGACTGTTCA AGCTGACATC ATACCCAGTC CAACAGACAC CAGCTGCCGT	1320
35	CAAGAAACTC CAGCCTTTCA AACTGACACC ACCCCCAGTG AGACAGGAGC CACTTCCACT	1380
	CCAGAAGCCA TCCCTGCTTT ATCTGAGACC ACCCCTACTG TGGTAGGAGC TGTATCTGCA	1440
40	CCGGCAGAAG CTAACACACC TCAGGATGCC ACATCTGCCC CAGAAGAGAC CAAGTAGCCA	1500
,,	AACTGTAGTC CTTCTAAAGG AGGACATGGC AGTCAAAAAG TCTGAGTAAA GCTGTTTTTT	1560
	GTATTTTATA TTTGCTTCTG CCATTTTACT GTCACTAATT AATGTTTAGT TCTTATATTT	1620
45	GTTAACTGAT TTCGGTGTCT TGAATATATT TTTTTAAATT ATGTGTATGA ACAATTCTAG	1680
	TITCATITGT TCAATCAGAA GAGCAAATAA CCATTCCTTT CATGTTTTGA TCACTGAGTG	1740
50	TGTCTGTAAT CATACCTACA TTAAAATCAT TTTCTATGAA TATATAATAT ATACTTCACA	1800
	TTTTTAGTGA ACTTCTCTAA AGAAGAGGAC AGAATATACT GGACTTAACC ACGAATACCC	1860
	TTGAGTGTCC AAATTGGGAA GGAACTKGTT TCTTCYGTTA TACTAYCAAA TGCTTAAATT	1920
55	CKGTTTCCTT TTTTCTTACC TTTGTTTGCT GTCTTTATGT AAAG	1964

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1522 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

	(XI)	, proomica .	DESCRIPTION	: SEQ ID NO	: 41:		
10	CGTGTCCGCG	CGCCTGGGAG	ACGCTGCCTC	GCCCGGACG	CGCCCGCGCC	CCCGCGGCTG	60
•	GAGGGTGGTC	GCCACTGGGA	CACTGTGAAC	CAGGAGTRAG	TCGGAGCTGC	CGCGCTGCCC	120
15	AGGCCATGGA	CTGTGAGGTC	AACAACGGTT	CCAGCCTCAG	GGATGAGTGC	ATCACAAACC	180
13	TACTGGTGTT	TGGCTTCCTC	CAAAGCTGTT	CTGACAACAG	CTTCCGCAGA	GAGCTGGACG	240
	CACTGGGCCA	CGAGCTGCCA	GTGCTGGCTC	CCCAGTGGGA	GGGCTACGAT	GAGCTGCAGA	300
20	CTGATGGCAA	CCGCAGCAGC	CACTCCCGCT	TGGGAAGAAT	AGAGGCAGAT	TCTGAAAGTC	360
	AAGAAGACAT	CATCCGGAAT	ATTGCCAGGC	ACCTCGCCCA	GGTCGGGGAC	AGCATGGACC	420
25	GTAGCATCCC	TCCGGGCCTG	GTGAACGGCC	TGGCCCTGCA	GCTCAGGAAC	ACCAGCCGGT	480
	CGGAGGAGGA	CCGGAACAGG	GACCTGGCCA	CTGCCCTGGA	GCAGCTGCTG	CAGGCCTACC	540
	CTAGAGACAT	GGAGAAGGAG	AAGACCATGC	TGGTGCTGGC	CCTGCTGCTG	GCCAAGAAGG	600
30	TGGCCAGTCA	CACGCCGTCC	TTGCTCCGTG	ATGTCTTTCA	CACAACAGTG	AATTTTATTA	660
	ACCAGAACCT	ACGCACCTAC	GTGAGGAGCT	TAGCCAGAAA	TGGGATGGAC	TGAACGGACA	720
35	GTTCCAGAAG	TGTGACTGGC	TAAAGCTCGA	TGTGGTCACA	GCTGTATAGC	TGCTTCCAGT	780
	GTAGACGGAG	CCCTGGCATG	TCAACAGCGT	TCCTAGAGAA	GACAGGCTGG	AAGATAGCTG	840
	TGACTTCTAT	TTTAAAGACA	ATGTTAAACT	TATAACCCAC	TTTAAAATAT	CTACATTAAT	900
40	ATACTTGAAT	GAAAATGTCC	ATTTACACGT	ATTTGAATGG	CCTTCATATC	ATCCACACAT	960
	GAATCTGCAC	ATCTGTAAAT	CTACACACGG	TGCCTTTATT	TCCACTGTGC	AGGTTCCCAC	1020
45	TTAAAAATTA	AATTGGAAAG	CAGGTTTCAA	GGAAGTAGAA	ACAAAATACA	ATTTTTTTGG	1080
	ТАААААААА	TTACTGTTTA	TTAAAGTACA	ACCATAGAGG	ATGGTCTTAC	AGCAGGCAGT	1140
	ATCCTGTTTG	AGGAAAGCAA	GAATCAGAGA	AGGAACATAC	CCCTTACAAA	TGAAAAATTC	1200
50	CACTCAAAAT	AGGGACTATC	YATCTTAATA	CTAAGGAACC	AACAATCTTC	CTGTTTAAAA	1260
	AACCACATGG	CACAGAGATT	CNGAACTAAA	GTGCTGCACT	CAAATGATGG	GAAGTCCCGG	1320
55	CCCCAGTACA	CCAGGGGCTT	TGGACTTTTT	TCAACTTCGT	TTCCTTTTGT	TTGGANTCCA	1380
	AAAGAACCAC	TTTGTGGTTC	TTAAAAGGGT	GTGAAGGTGA	TTTAAGGGGC	CCAGGTCAGC	1440
	CACTGGTTGG	TTTACAAAAT	CNGGGTAACT	AACTGCATAC	AACTITTTCC	CNTTTCCATG	1500
60	NCATCAGGAC	TTTGCTAAAG	AC				1522

5	(2) INFORMATION FOR SEQ ID NO: 42:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 875 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:	
15	TGGGATTTCC CTTTATCATG GAGGCCTTGT CCCACTTCCT CTATGTCCCT TTCCTTGGTG	60
	TCTGTGTCTG TGGGGCCATC TACACTGGCC TGTTCCTTCC TGAGACCAAA GGCAAGACCT	120
20	TCCAAGAGAT CTCCGAGGAA TTACACAGAC TCAACTTCCC CAGGCGGGCC CAGGGCCCCA	180
20	CGTGGAGGAG CCTGGAGGTT ATCCAGTCAA CAGAACTCTA GTCCCAAAGG GGTGGCCGTA	240
	GCCAAAGCCA GCTACCGTCC TGTCCTCTGC TTCCTGCCAG GGCCCTGGTC CTCAMTYCCT	300
25	YCTGCATTCC TCATTTAAGG AGTGTTTATT GAGCACCCTT TGTGTGCAGA CATGGCTCCA	360
	GGTGCTTAGC AATCAWTGGT GAGCGTGGTA TCCAGGCTAA AGGTAATTAA CTGACAGRAA	420
30	ATCAGTAACA ACATAATTAC AGGYTGGTTG TGGCAGYTCA TGACTGTAAT CCCAGCACTT	480
30	TTGGGAGCCA AGGTGGGARG ATCAATTGAG GCCAGAGTTT GAAAMCAGCT AGGTAACATA	540
	GTGAGACCCC CTATCTCTAC AAAAAATTTT AAACATTAGC TGGGCATGGT GGTATGTGCT	600
35	AACAGCTCTA GCTACTCAGG AGGCTGAGGC AGCAGGATCA CTTGAGTCCA AGAGTTCAAG	660
	GTAGCAGTAA GCTACAATCA CACCACTGCA TGCCAGACTG GGTGACAGAG GGAGACTTCA	720
40	TCTCTTTAAA ACATAATAAT AATAATTACA GACTCAGGAA ATGCAGTGAA AGAAAAATAC	780
40	AGGTTGGCCA GGTGAGGTGG CTGATGCCTG TAATCCCAGC ACTTTGGGAG GCCAAGATGG	840
	GAAGATTGCT TTGAGACCAG AAGTTTGAGA CCAGC	875
45		
	(2) THEODY MICH FOR SEC ID NO. 42.	
50	(2) INFORMATION FOR SEQ ID NO: 43:	
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 843 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
55	(D) TOPOLOGY: linear	
_ -	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:	
	CCCACGCGGT CCGNATCGTC CTTCCCTCAC TTCAGAGGGT GGCCAGAGCT GAATACCCAG	60
60	ACACCCACAA CTAACCCTTC ACTTCCAAAA CATCATCACC ATCTATCATC CCACCTCTCT	120

	CACCIGACAG TIACAGAGGA AACCCGCACC CAGAATGCAC GIGCTGTCTT ATGGGAACAC	180
5	TCAGCGCAGA GTGCTCAGGT CCGGCCACAC TCGGGCTGTG CTTGGTCGTG CCATGGAATT	240
,	CCTCAGGACT TTCTCAGCCT CCCTAATGGC AGAAGCCCCT TTACAGCAAG ACATTTACCG	300
	TTTGTCTGAA AATAGCCGAA CTGAGCTTTT CTTCAGGCTA TATGAGAAGT CTCTAGACAG	360
10	TGGGCACCGT CAGAAAGCCC AGAGCCTTGT GATAGCTCCC ACCCTGCCTG GCTCAGATCT	420
	TCCCATTTTT TTTCCTCTGG CACTAACCTC ACCTTTTGTT TTTTTTGTGTT TGTGTTTGTT	480
15	TTTGTTTTTG CAGAGTTGGA TTACAGAAAC TCCTATGAAA TTGAATATAT GGAGAAAATT	540
	GGCTCCTCCT TACCTGTAAG TTCGTCTGCC TCGGGCCACT TAGGGGACTC GCTTTCCTGC	600
	CTTCAGGGGC CTCCTCCCCT GTGCAGAGTG TCTCTGGGAG CTCAGACCCC AAATCGAGTG	660
20	TTTTCTGTGT ACACAGCTTC CCGGGTGCAC AGCAATGATG GACTGGGGCT GGGGGGTTGA	720
	GGTTTGTACT CAATCCACTT CGTTTGACAT TTTCAGGGAG AAAATGATAG AATACAATŢA	780
25	GACGTCCTGC AGAATTACTT TCCTAGACTG AGAAAGAGCT AGAGATTTCT TTAAAAAAAA	840
	AAA	843
30	(2) INFORMATION FOR SEQ ID NO: 44:	
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 489 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:	
	CTCTTAGGCT TTGAAGCATT TTTGTCTGTG CTCCCTGATC TTCAGGTCAC CACCATGAAG	60
	TTCTTAGCAG TCCTGGTACT CTTGGGAGTT TCCATCTTTC TGGTCTCTGC CCAGAATCCG	120
45	ACAACAGCTG CTCCAGCTGA CACGTATCCA GCTACTGGTC CTGCTGATGA TGAAGCCCCT	180
	GATGCTGAAA CCACTGCTGC TGCAACCACT GCGACCACTG CTGCTCCTAC CACTGCAACC	240
50	ACCGCTGCTT CTACCACTGC TCGTAAAGAC ATTCCAGTTT TACCCAAATG GGTTGGGGAT	300
- 0	CTCCCGAATG GTAGAGTGTG TCCCTGAGAT GGAATCAGCT TGAGTCTTCT GCAATTGGTC	360
	ACAACTATTC ATGCTTCCTG TGATTTCATC CAACTACTTA CCTTGCCTAC GATATCCCCT	420
55	TTATCTCTAA TCAGTTTATT TTCTTTCAAA TAAAAAATAA CTATGAGCAA CAAAAAAAA	480
	ААААААА	489

	(2) INFORMATION FOR SEQ ID NO: 45:	
5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 534 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 45:	
	GAAGCAGTGT GTATCTATGA TTATATCTCT GTTCATCTAT ATATTTTTGA CATGTAGCAA	60
15	CACCTCTCCA TCTTATCAAG GAACTCAACT CGGTCTGGGT CTCCCCAGTG CCCAGTGGTG	120
13	GCCTTTGACA GGTAGGAGGA TGCAGTGCTG CAGGCTATTT TGTTTTTTGT TACAAAACTG	180
	TCTTTTCCCT TTTCCCCTCC ACCTGATTCA GCATGATCCC TGTGAGCTGG TTCTCACAAT	240
20	CTCCTGGGAC TGGGCTGAGG CAGGGGCTTC GCTCTATTCT CCCTAACCAT ACTGTCTTCC	300
	TTTCCCCTTG CCACTTAGCA GTTATCCCCC CAGCTATGCC TTCTCCCTCC CTCCCTTGCC	360
25	CTGGCATATA TTGTGCCTTA TTTATGCTGC AAATATAACA TTAAACTATC AAGTGAAAAA	420
23	AAAAAAAAA AAAACTCCAA GGGGGGCCG GTACCCAATT CCCCCTATAN TGAGTCNTAT	480
	TACAATTCAC TGGGCCGTCG TTTTACAACG TCGTGAATGG GAAAACCTGG GCGT	534
30		
	(2) INFORMATION FOR SEQ ID NO: 46:	
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1374 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double	
40	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:	
	GGCACGAGTC CGGGATGAGC TCAGCCGCGG CCGACCACTG GGCGTGGTTG CTGGTGCTCA	60
45	GCTTCGTGTT TGGATGCAAT GTTCTTAGGA TCCTCCTCCC GTCCTTCTCA TCCTTCATGT	120
	CCAGGGTGCT GCAGAAGGAC GCGGAGCAGG AGTCACAGAT GAGAGCGGAG ATCCAGGACA	180
50	TGAAGCAGGA GCTCTCCACA GTCAACATGA TGGACGAGTT TGCCAGATAT GCCAGGCTGG	240
	AAAGAAAGAT CAACAAGATG ACGGATAAGC TCAAAACCCA TGTGAAAGCT CGGACAGCTC	300
	AATTAGCCAA GATAAAATGG GTGATAAGTG TCGCTTTCTA CGTATTGCAG GCTGCCCTGA	360
55	TGATCTCACT CATTTGGAAG TATTATTCTG TCCCTGTGGC TGTCGTGCCG AGTAAATGGA	420

TGATCTCACT CATTTGGAAG TATTATTCTG TCCCTGTGGC TGTCGTGCCG AGTAAATGGA

TAACCCCTCT AGACCGCCTG GTAGCCTTTC CTACTAGAGT AGCAGGTGGT GTTGGAATTA

CCTGTTGGAT TTTAGTCTGT AACAAAGTTG TCGCTATTGT GCTTCATCCG TTCAGCTGAA

480

596

	CAGGAGGATG GATACAGCCG CGAGGCTAAA AAACGGATTT CCTCTTCCTA GCTTAAAATC	600
	TGATTTACAC TGTTTTGTTT TTTAAGAAAC AAAAGTGCAT AGTTTAGATT TTTTTTTTTG	660
5	TTGAATATGT TTGTTCTTGG ACTTTATGAG AGAGTCTTAT AAGAATCACG ATTTTCTACA	720
	CCTGTCATTG AGCCAAGAAA GTCCAGTTTA TGACACGTAT GTACTAGTGA ACACCGTCCT	780
10	CGATCTGTAC GAAATGTGAA ATGTTTAGGG ACATCTCCAT GCTGTCACTT GTGATTTGCC	840
10	CTCTTATGTA TTTTGGTCAT ATTGCCAACT GGAAAGTCAA AATTTTCTAA CAACTTTAAG	900
	TAAGTTCTTT GAAGACTTAG TGCTGTTTTT AATCCAGTTT AGAAAGTAAC TTAATTTTAA	960
15	TACCACTACT AAAAATTCGA AAATTTCTTC TTTAATCACA TTCAATATGG TTAAAAGAAC	1020
	AACACTAATT GACATTGCGT GGGCTTTTTC TCCCTTTGTT TAAAATGTCA TTTGTTGAGC	1080
20	AAGAGTTGTA TAGTATTATC TACTTACTTG AGGCTGTTAA TTTTTCATTA CAGTGTTTTG	1140
20	TAAATGTATC CACGAGACCA TGATGCATTG TTTTGTGCTC AACTTGTGTT TTGTATTTAA	1200
	AGCATTTTGA ATGAAGTGTA TTTTATAAGC ATTTAATATT TATGCTCTTT AGAATGGAAC	1260
25	ACAGAAAACA AACCTTATAA GTCCTGATTA ATCTGAACCA ATAACCTGTG TGGCCTACAA	1320
	AGTATAATTC TATTAAATGT TCCTTAAAAC AAAAAAAAAA	1374
30		
	(2) INFORMATION FOR SEQ ID NO: 47:	
	(i) SEQUENCE CHARACTERISTICS:	
35	(A) LENGTH: 596 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:	
	GAATTCGNCA CGAGATTACT TGGACATGAA AGAACTCAGG TTCAAGTITA TTCATTTACT	60
	AAGTTAGTTA AATCATGTGC CTTCCATGAG CCTTCATTTG GTAACTTGGA AAATGGAAAT	120
45	AATAACACTA GTCATATATA TTCTACACTG CTACCATATG GACCAAAGGG ATTATAGATT	180
	ACAATCACCA TCATTCCTGC TGACAGGTAT ATAGAAAACA ATTTCATTGA AGAAAAGTCC	240
50	TTACATTTAT CCTTTTCCTA ATATCTGCAT GGGTAAACTA ATAAATATAG TCATTAGAAA	300
	ACCCTTATTA TTATTATTAG TTCAATGTGA GAACTGCTGC AGAAAAAATA TGCTTTATAA	366
	TATTTTCTTG AATATACATA ATATTCATAA ATTTTCAAAT CATTGAAAAT TACCTTAAAA	420
55	TTGGAAAAAA TGTGCATTTC TACTCATATA ACAGTATAAA ATTCCTATGT CAATCTCTTT	48
	TTTTTTTTT TGTTTTGAGT TGGAGTCTCG CTCTGTCGCC CAGGCTGGGC AACAGAGCAG	54

GACCCTGTCT TAATTAAAAA AAAAAAAAA AAACTCGAGG GGGGCCCGGT ACCCTA

5	(2) INFORMATION FOR SEQ ID NO: 48:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 851 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:	
15	CACATGAAGA CACACAGTGG TGAGAAGCCC TTCCGCTGCG CCCGCTGTCC TTATGCCTCT	60
	CCTCATCTGG ATAACCTGAA ACGGCACCAG CGCGTCCATA CAGGAGAGAA GCCCTACAAG	120
20	TGCCCCCTCT GCCCTTATGC CTGTGGCAAT CTGGCCAACC TCAAGCGTCA TGGTCGCATC	180
20	CACTCTGGTG ACAAACCTTT TCGGTGTAGC CTTTGCAACT ACAGCTGCAA CCAGAGCATG	240
	AACCTCAAAC GTCACATGCT GCGGCACACA GGCGAGAAGC CTTCCGCTGT GCCACCTGCG	300
25	CCTATACCAC GGGCCACTGG GACAACTACA AGCGCCACCA GAAGGTGCAT GGCCACGGTG	360
	GGGCAGGAGG GCCTGGTCTC TCTGCCTCTG AGGGCTGGGC CCCACCTCAT AGCCCACCCT	420
30	CTGTTTTGAG CTCTCGGGGC CCACCAGCCC TGGGGACTGC TGGCAGCCGG GCTGTCCACA	480
50	CAGACTCATC CTGAACTAGG TCCTTCTTCC CCATGTTTTA TACAGACGGA CCAGAAGCCA	540
	CCTTTTTCTC CCCCGCTGGC CAGGGGCTCC ACACAGACTA ACGTAGGCAC TATAAGGACC	600
35	AGCCCAACCC CATGGGCGGG GGGGCCCATA TGGACCAGGG GACCTTGCCT TGACTGAGGC	660
	ACTTCACGAG CTCAGTGAGA AGGGCCCTGT ATTCACCTCC ACTGCCCCCA GGGGCTGTGG	720
40	ACAAACCGGC TGGGGGACTG CCCAGCCTCC CACCTGTTTA TTTAACTTAT TTCAGTGCTT	780
40	TATAATAAAG GAAACACTAA CAAAGCCATG TCTATGCTGA ATTGGCAATG GCAGGCAATT	840
	TGGCCTTACC C	851
45		
	(2) INFORMATION FOR SEQ ID NO: 49:	
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2020 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 49:	
	GTGAAATGAA AACAGTCTTT TTATAGCCTT TAGCTTGTGA GTTTGGAAGT TTGGGGGGTC	60
60	TTATGTTTGT TTTGCCTCTT CTGTTTCTTG GAGGAGAGTT GAGGCTTTTC TTAGGTGCAT	120

	ACACAGACCC	AGGTGAACAC	GCTGACTGTG	AACCTGCCCT	GTATCCGGAG	CTGTGCTGGG	180
5	CACTGAGGGG	ATGCAACAAA	ATTAGGAGAG	GWTCCTTGCT	CCCAACGTCT	ACTTCTCCTA	240
3	CCTCAACAGG	GGTCCAGGGT	GCAGTGAACT	CAGTTCTTGG	CCCTTGGGTG	AGGATTCATG	300
	GATGAATGAA	AGCTAGACCT	GATGGGGAGG	CATTATGACT	AAATAGGCCC	AGCCTCCTTC	360
0	CCTTCCAGCT	CTGTCCTAGG	AGCATAGGCG	GGAAATCTGA	GTAGAGTCTG	ACTGCAGTTT	420
	TTGCTTATGA	TTTGTAAAAG	CCGTCATGGG	GTCAATAAGA	AAATAGGGGT	GATGGAGGG	480
15	GAGAAGCCCA	GGACTGGGAG	AATCGCACGT	GCCCCAGGGG	TTTTCACCAA	GGATTTTCAA	540
	GACAAACTGG	AGTAAGAATT	AAAGCCCCAG	AGGATTTAAT	TATCCTGGTT	TGCAAAAGAG	600
	CCTCCCATGC	CAGTACCGCC	CAGCCTTGGA	GGCCGGAATG	CTCATGGCCC	CTGTGGTCTG	660
20	CTTGTCCTTC	AGCCCATGCC	CAGCAGATAC	CTCTCTGACT	GGAGACGGGC	TCAAAGCTGG	720
	ATTAGAAAGG	GGAGMGGCAC	TIGIGACTIT	GTTTGACTCT	GTGACTCACT	TCCTCGCTCA	780
25	CACCTTGTTT	GAACTACTGG	ACTITICAACT	GGCTTTCCTT	AGGTCAGGCA	AGCAGACAGC	840
-3	TCCCCACTGA	AGAGGTCTGT	ACAGTGACAA	ccceecce	CAGCAAGGAC	ACAGATGCAG	900
	CCACAGTAAG	GCTCCATCAG	GACTGGGTCA	GTGATGGCAA	CAGGATGGCC	AAGGATGGCT	960
30	CTAGAACAYT	CTGTCCATGC	GTCACTCCCC	CCAGTTTTRT	TTTTAGCTTT	GGCTTCAGGG	1020
	AGTGACAGCC	ATCACAAATA	GCCACATTCT	GCTCTACTCT	CCAACATACC	AGATTSTACA	1080
35	CTGTTGTTAT	TTCATGAGAC	GTGAATGTTG	CAGAGAGTGG	GGGGATTCTG	GTTGTTAAGG	1140
	AACTTACACT	GGGGAGCTTT	ACTOTTCCGT	GTCAACAATG	TGACTACATG	TTCTCCAGAT	1200
	TAGCCACACA	TGCAAACATC	AGTGTCCTTC	TAGCTTTANC	CGAGAAAGAA	ACCAGTCCCA	1260
40	GGGAATGAAT	GGTGGTCTCC	CCACTCCCGG	CAGCACTITA	. GGCAGCCCAT	AAGCTATGCG	1320
	AGAATGTGAA	CGCTCACCTT	GCTCCGTCAC	GGTTCTGACC	TACCACATAA	ACAGGAAGAA	1380
45	GCCAGTGACC	GGAACAGCTC	TAGGAATAAC	AAGTCAGAAT	AGAAGTGTCC	TTTATATTAC	1440
	CAGAAAATAT	GGGCTTGGCC	TAAGTCGCTG	TCTCCTAACC	TGCCGGGGTC	ATTCCCCACC	1500
	AAACACCCCA	TACTAAGGAG	CCATGAGCCA	CCTGGACATI	CACCITITCI	TTGACCATCT	1560
50	GGAGTCTGGG	GCAACTTAAG	GAAGGCNCCA	CACAGTGGTC	CAGGCACATI	TCCAAGCGTA	1620
	GGTGTCCCTG	GCTTTTGTGG	CCAAAGCTAG	TGTTATGGTC	: AACAACAGGC	CAGGGTCTGT	1680
55	GGGGCACTGA	CCTTGAAAGT	GGCAAAATGG	AGGTTTCACA	GGCTGTGCGG	GAGCAGGACG	1740
- •	GCTTGCTTCA	TCTAACAATC	TCAGTTTCCT	CAAAAAATT	AAAGAAAGGA	AAAGATTTCA	1800
	TAAGCAGGTG	TCAGTGGACA	GTTTAAGYAC	TTAACCATT	CICTITCIA	TTATGGATGT	1860
60	GAACTCTCCT	TAAATAAAAT	יירבידידידים י	ייבעניע עיבענייבעני	ר קייייים איי	י יים ארא כייייאיי	1920

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	TAAGTCGGTT GTGTATATGT GTAACTAATG TAACTGCCTT TTAAAATTTC ATTACAATAA	1980
5	AAATGACTTT GCTCTGAAMA AAAAAAAAA AAAAACTCGA	2020
J		
	(2) INFORMATION FOR SEQ ID NO: 50:	
10	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2432 base pairs (B) TYPE: nucleic acid	
15	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 50:	
	ATGAAGGGTC GTTGGTGGGA AAGATGGCGG CGACTCTGGG ACCCCTTGGG TCGTGGCAGC	60
20	AGTGGCGGCG ATGTTTGTCG GCTCGGGATG GGTCCAGGAT GTTACTCCTT CTTCTTTTGT	
		120
25	TGGGGTCTGG GCAGGGGCCA CAGCAAGTCG GGGCGGGTCA AACGTTCGAG TACTTGAAAC	180
25	GGGAGCACTC GCTGTCGAAG CCCTACCAGG GTGTGGGCAC AGGCAGTTCC TCACTGTGGA	240
	ATCTGATGGG CAATGCCATG GTGATGACCC AGTATATCCG CCTTACCCCA GATATGCAAA	300
30	GTAAACAGGG TGCCTTGTGG AACCGGGTGC CATGTTTCCT GAGAGACTGG GAGTTGCAGG	360
	TGCACTTCAA AATCCATGGA CAAGGAAAGA AGAATCTGCA TGGGGATGGC TTGGCAATCT	420
	GGTACACAAG GAATCGGATG CAGCCAGGGC CTGTGTTTGG AAACATGGAC AAATTTGTGG	480
35	GGCTGGGAGT ATTTGTAGAC ACCTACCCCA ATGAGGAGAA GCAGCAAGAG CGGGTATTCC	540
	CCTACATCTC AGCCATGGTG AACAACGGCT CCCTCAGCTA TGATCATGAG CGGGATGGGC	600
40	GGCCTACAGA GCTGGGAGGC TGCACAGCCA TTGTCCGCAA TCTTCATTAC GACACCTTCC	660
+0	TGGTGATTCG CTACGTCAAG AGGCATTTGA CGATAATGAT GGATATTGAT GGCAAGCATG	720
	AGTGGAGGGA CTGCATTGAA GTGCCCGGAG TCCGCCTGCC CCGCGGCTAC TACTTCGGCA	780
45	CCTCCTCCAT CACTGGGGAT CTCTCAGATA ATCATGATGT CATTTCCTTG AAGTTGTTTG	840
	AACTGACAGT GGAGAGAACC CCAGAAGAGG AAAAGCTCCA TCGAGATGTG TTCTTGCCCT	900
	CAGTGGACAA TATGAAGCTG CCTGAGATGA CAGCTCCACT GCCGCCCCTG AGTGGCCTGG	960
50	CCCTCTTCCT CATCGTCTTT TTCTCCCTGG TGTTTTCTGT ATTTGCCATA GTCATTGGTA	1020
	TCATACTCTA CAACAAATGG CAGGAACAGA GCCGAAAGCG CTTCTACTGA GCCCTCCTGC	1080
55	TGCCACCACT TTTGTGACTG TCACCCATGA GGTATGGAAG GAGCAGGCAC TGGCCTGAGC	1140
	ATGCAGCCTG GAGAGTGTTC TTGTCTCTAG CAGCTGGTTG GGGACTATAT TCTGTCACTG	1200
	GAGTITIGAA TGCAGGGACC CCGCATTCCC ATGGTTGTGC ATGGGGACAT CTAACTCTGG	1260

240

	TCTGGGAAGC CACCCACCCC AGGGCAATGC TGCTGTGATG TGCCTTTCCC	TGCAGTCCTT	1320
	CCATGTGGGA GCAGAGGTGT GAAGAGAATT TACGTGGTTG TGATGCCAAA	ATCACAGAAC	1380
5	AGAATTTCAT AGCCCAGGCT GCCGTGTTGT TTGACTCAGA AGGCCCTTCT	ACTTCAGTTT	1440
	TGAATCCACA AAGAATTAAA AACTGGTAAC ACCACAGGCT TTCTGACCAT	CCATTCGTTG	1500
10	GGTTTTGCAT TTGACCCAAC CCTCTGCCTA CCTGAGGAGC TTTCTTTGGA	AACCAGGATG	1560
	GAAACTICTT CCCTGCCTTA CCTTCCTTTC ACTCCATTCA TTGTCCTCTC	TGTGTGCAAC	1620
	CTGAGCTGGG AAAGGCATTT GGATGCCTCT CTGTTGGGGC CTGGGGCTGC	AGAACACACC	1680
15	TGCGTTTCAC TGGCCTTCAT TAGGTGGCCC TAGGGAGATG GCTTTCTGCT	TTGGATCACT	1740
	GTTCCCTAGC ATGGGTCTTG GGTCTATTGG CATGTCCATG GCCTTCCCA	TCAAGTCTCT	1800
20	TCAGGCCCTC AGTGAAGTTT GGCTAAAGGT TGGTGTAAAA ATCAAGAGA	GCCTGGAAGA	1860
	CATCATGGAT GCCATGGATT AGCTGTGCAA CTGACCAGCT CCAGGTTTGA	TCAAACCAAA	1920
	AGCAACATTT GTCATGTGGT CTGACCATGT GGAGATGTTT CTGGACTTGC	TAGAGCCTGC	1980
25	TTAGCTGCAT GTTTTGTAGT TACGATTTTT GGAATCCCAC TTTGAGTGC	GAAAGTGTAA	2040
	GGAAGCTTTC TTCTTACACC TTGGGCTTGG ATATTGCCCA GAGAAGAAA	TIGGCTITIT	2100
30	TTTTCTTAAT GGACAAGAGA CAGTTGCTGT TCTCATGTTC CAAGTCTGAG	G AGCAACAGAC	2160
	CCTCATCATC TGTGCCTGGA AGAGTTCACT GTCATTGAGC AGCACAGCC	r GAGTGCTGGC	2220
	CTCTGTCAAC CCTTATTCCA CTGCCTTATT TGACAAGGGG TTACATGCT	G CTCACCTTAC	2280
35	TGCCCTGGGA TTAAATCAGT TACAGGCCAG AGTCTCCTTG GAGGGCCTG	G AACTCTGAGT	2340
	CCTCCTATGA ACCTCTGTAG CCTAAATGAA ATTCTTAAAA TCACCGATG	G AACCAAAAA	2400
40	AA AAAAAAAAA AAAAAAAAA AAAAAAAAA		2432
	(2) INFORMATION FOR SEQ ID NO: 51:		
45	(i) SEQUENCE CHARACTERISTICS:		
	(A) LENGTH: 2340 base pairs(B) TYPE: nucleic acid		
50	(C) STRANDEDNESS: double (D) TOPOLOGY: linear		
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 51:		
~ ~	GACGCTGGGG GCGGGTGGGG GCGCGGGTA CCGGGCTGGA CGGCCGGCC	G GCGCCCCCTC	60
55	ATTAGTATGC GGACGAAGCG GCGGGCTGCG CGGAGNGACG TCCCCTGCA	G CCGCGGACCG	120
	AGGCAGCGGC GGCACCTGCC GGCCGAGCAA TGCCAAGTGA GTACACCTA	T GTRAAACTGA	180

GAAGTGATTG CTCGAGGCCT TCCCTGCAAT GGTACACCCG AGCTCAAAGC AAGATGAGAA

	GGCCCAGCTT GTTATTAAAA GACATCCTCA AATGTACATT GCTTGTGTTT GGAGTGTGGA	300
5	TCCTTTATAT CCTCAAGTTA AATTATACTA CTGAAGAATG TGACATGAAA AAAATGCATT	360
3	ATGTGGACCC TGACCATGTA AAGAGAGCTC AGAAATATGC TCAGCAAGTC TTGCAGAAGG	420
	AATGTCGTCC CAAGTTTGCC AAGACATCAA TGGCGCTGTT ATTTGAGCAC AGGTATAGCG	480
10	TGGACTTACT CCCTTTTGTG CAGAAGGSCC CCAAAGACAG TGAAGCTGAG TCCAAGTACG	540
	ATCCTCCTTT TGGGTTCCGG AAGTTCTCCA GTAAAGTCCA GACCCTCTTG GAACTCTTGC	600
15	CAGAGCACGA CCTCCCTGAA CACTTGAAAG CCAAGACCTG TCGGCGCTGT GTGGTTATTG	660
15	GAAGCGGAGG AATACTGCAC GGATTAGAAC TGGGCCACAC CCTGAACCAG TTCGATGTTG	720
	TGATAAGGTT AAACAGTGCA CCAGTTGAGG GATATTCAGA ACATGTTGGA AATAAAACTA	780
20	CTATAAGGAT GACTTATCCA GAGGGCGCAC CACTGTCTGA CCTTGAATAT TATTCCAATG	840
	ACTTATTTGT TGCTGTTTTA TTTAAGAGTG TTGATTTCAA CTGGCTTCAA GCAATGGTAA	900
25	AAAAGGAAAC CCIGCCATTC TGGGTACGAC TCTTCTTTTG GAAGCAGGTG GCAGAAAAA	960
	TCCCACTGCA GCCAAAACAT TTCAGGATTT TGAATCCAGT TATCATCAAA GAGACTGCCT	1020
	TTGRACATCC TTCAGTACTC AGAGCCTCAG TCAAGGTTCT GGGGGCCGAG ATAAGAACGT	1080
30	CCCCACAATC GGTGTCATTG CCGTTGTCTT AGCCACACAT CTGTGCGATG AAGTCAGTTT	1140
	GGCGGGTTTT GGATATGACC TCAATCAACC CAGAACACCT TTGCACTACT TCGACAGTCA	1200
35	ATGCATGGCT GCTATGAACT TTCAGACCAT GCATAATGTG ACAACGGAAA CCAAGTTCCT	1260
	CTTAAAGCTG GTCAAAGAGG GAGTGGTGAA AGATCTCAGT GGAGGCATTG ATCGTGAATT	1320
	TTGAACACAG AAAACCTCAG TTGAAAATGC AACTCTAACT CTGAGAGCTG TTTTTGACAG	1380
40	CCTTCTTGAT GTATTTCTCC ATCCTGCAGA TACTTTGAAG TGCAGCTCAT GTTTTTAACT	1440
	TTTAATTTAA AAACACAAAA AAAATTTTAG CTCTTCCCAC TTTTTTTTTC CTATTTATTT	1500
45	GAGGTCAGTG TTTGTTTTTG CACACCATTT TGTAAATGAA ACTTAAGAAT TGAATTGGAA	1560
	AGACTTCTCA AAGAGAATTG TATGTAACGA TGTTGTWFTG ATTTTTAAGA AAGTAATTTA	1620
	ATTTGTAAAA CTTCTGCTCG TTTACACTGC ACATTGAATA CAGGTAACTA ATTGGAAGGA	1680
50	GAGGGGAGGT CACTCTTTTG ATGGTGGCCC TGAACCTCAT TCTGGTTCCC TGCTGCGCTG	1740
	CTTGGTGTGA CCCACGGAGG ATCCACTCCC AGGATGACGT GCTCCGTAGC TCTGCTGCTG	1800
55	ATACTGGGTC TGCGATGCAG CGGCGTGAGG CCTGGGCTGG TTGGAGAAGG TCACAACCCT	1860
	TCTCTGTTGG TCTGCCTTCT GCTGAAAGAC TCGAGAACCA ACCAGGGAAG CTGTCCTGGA	1920
	GGTCCCTGGT CGGAGAGGGA CATAGAATCT GTGACCTCTG ACAACTGTGA AGCCACCCTG	1980
60	GGCTACAGAA ACCACAGTCT TCCCAGCAAT TATTACAATT CTTGAATTCC TTGGGGATTT	2040

	TTTACTGCCC TTTCAAAGCA CTTAAGTGTT AGATCTAACG TGTTCCAGTG TCTGTCTGAG	2100
5	GTGACTTAAA AAATCAGAAC AAAACTTCTA TTATCCAGAG TCATGGGAGA GTACACCCTT	2160
J	TCCAGGAATA ATGTTTTGGG AAACACTGAA ATGAAATCTT CCCAGTATTA TAAATTGTGT	2220
	ATTTAAAAAA AAGAAACTTT TCTGAATGCC TACTGGCGGT GTATACCAGG CAGTGTGCCA	2280
10	GTTTAAAAAG ATGAAAAAGA ATAAAAACTT TTGAGGAAMA AAAAAAAAAA AAAAACTCGA	2340
15	(2) INFORMATION FOR SEQ ID NO: 52:	
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 601 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 52:	
25	AGTAGGGGAG ACTGAGACTG ACCGGTAGCC AGGCAGGCGG ACGACGCACG CCCGGACAGA	60
	CTGAGCAGGC GCCGGAGAAC CACTCACAGG TTCCCCCCGC CTTTCCCTTT GAAANCTAGG	120
30	CTTTTGCCTT TCCCGTGGCG CCCGAGAGAG AATGCTGGAC TCTGCCGACT TCAGCGCAAC	180
	TAANGATTTC TCAAGCTAGG GGACAAACGA TCAGCCCAAT CCTGAGAAGG GGGGAACCAA	240
	GCACCCCGTC CCCATCCCCC TCCCCTCCCC CGACTAAACT CGGGCGCCAA ACCCAGCCCT	300
35	TCTCTAACCA CCCTACTTCC TCCTCTCTT TCTAGCATGG TGGCTGTATG GACAGTCTGA	360
	CAGAACAGAG ACTGACATCT CCCAATCTGC CGGCCCCCCA CCTGGAACAC TACÁGTGTTC	420
40	TGCATTGCAC CATGACCCTG GATGTGCAAA CTGTAGTCGT TTTTGCCGTG ATTGTAGTCC	480
	TCCTGCTTGT CAATGTCATA CTCATGTTTT TCCTGGGAAC GCGCTGAATG GAGTCCAGNC	540
	ACCTGAGCTG TCGCGAACTC TCGCTTTGAT TTCATCCCGA GAGCCACCGA GAAGAAAAAA	600
45	A	601
50	(2) INFORMATION FOR SEQ ID NO: 53:	
	(i) SEQUENCE CHARACTERISTICS:	
55	(A) LENGTH: 359 base pairs (B) TYPE: nucleic acid	
55	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:	
60	CTCGTGCCGA ATTCGGCACG AGAGATGGTA CTTTTAAGAG GTAATTAGGT TGCTAAGATG	60

	GATTAACATC	TTTCTCTTGA	CACTGAGACT	GGGTTCTCCT	GGGAATGGTT	AGTTCCCAAG	120
5	AGAGTGAGTT	GTTATAAAAC	AATGCTGCCT	CTTCTATTTT	GCGCTTTTTG	TTTGCACAAA	180
,	CTCGGTCCCC	TTCTGTTTCT	CTACGATGTT	TTGATGCRGC	ATGAGGCAGT	CATGAGAACC	240
	CACCAGATAC	AGCTGCCTGA	TCCTGAATTT	CCCAGCCAAC	AGAACCAAGT	ССТАААТАА А	300
10	ACTCTTTTTA	ATAAGTTAAA	АААААААА	АААААААА	AANAAANANA	AAAAAAA	359
15	(2) INFORMA	TION FOR SE	Q ID NO: 54	l:			
20	(i)	(B) TYPI (C) STR	HARACTERIST: GTH: 1141 b E: nucleic a ANDEDNESS: O DLOGY: line	ase pairs acid double	·		
	(xi)	SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 54:		
25	GGCACGAGCT	GCTGAGGCGT	GAGAATGGCG	TCCCGCGGCC	GGCGTCCGGA	GCATGGCGGA	60
	CCCCAGAGC	TGTTTTATGA	CGAGACAGAA	GCCCGGAAAT	ACGTTCGCAA	CTCACGGATG	120
30	ATTGATATCC	AGACCAGGAT	GGCTGGGCGA	GCATTGGAGC	TTCTTTATCT	GCCAGAGAAT	180
50	AAGCCCTGTT	ACCTGCTGGA	TATTGGCTGT	GGCACTGGGC	TGAGTGGAAG	TTATCTGTCA	240
	GATGAAGGGC	ACTATTGGGT	GGGCCTGGAT	ATCAGCCCTG	CCATGCTGGA	TGAGGCTGTG	300
35	GACCGAGAGA	TAGAGGGAGA	CCTGCTGCTG	GGGGATATGG	GCCAGGGCAT	CCCATTCAAG	360
	CCAGGCACAT	TTGATGGTTG	CATCAGCATT	TCTGCTGTGC	AGTGGCTCTG	TAATGCTAAC	420
40	AAGAAGTCTG	AAAACCCTGC	CAAGCGCCTG	TACTGCTTTT	TTGCTTCTCT	TTTTTCTGTT	480
. •	CTCGTCCGGG	GATCCCGAGC	TGTCCTGCAG	CTGTACCCTG	AGAACTCAGA	GCAGTTGGAG	540
•	CTGATCACAA	CCCAGGCCAC	AAAGGCAGGC	TTCTCCGGTG	GCATGGTGGT	AGACTACCCT	600
45	AACAGTGCCA	AAGCAAAGAA	ATTCTACCTC	TGCTTGTTTT	CTGGGCCTTC	GACCTTTATA	660
	CCAGAGGGGC	TGAGTGAAAA	TCAGGATGAA	GTTGAACCCA	GGGAGTCTGT	GTTCACCAAT	720
50	GAGAGGTTCC	CATTAAGGAT	GTCGAGGCGG	GGAATGGTGA	GGAAGAGTCG	GGCATGGGTG	780
	CTGGAGAAGA	AGGAGCGGCA	CAGGCGCCAG	GGCAGGGAAG	TCAGACCTGA	CACCCAGTAC	8,40
	ACCGGCCGCA	AGCGCAAGCC	CCGCTTCTAA	GTCACCACGC	GGTTCTGGAA	AGGCACTTGC	900
55	CTCTGCACTT	TTCTATATTG	TTCAGCTGAC	AAAGTAGTAT	TTTAGAAAAG	TTCTAAAGTT	960
	ATAAAAATGT	TTTCTGCAGT	AAAAAAAAG	TTCTCTGGGC	CGGCCTGGT	GGCTCACACC	1020
60	TGTAATCCCA	GCACCTTGGG	AGGCTGAGGT	GGGAGGATCA	TTTGAGGCCA	GGAGTTTGAG	1080

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1140 1141 Α 5 (2) INFORMATION FOR SEQ ID NO: 55: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1560 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 55: TCCTTCTCTG GGGCGTCGC GTTGGCAGCG GATGCGGGAA GCCGGACTCT GGGCGTCATG 60 20 TACTACAAGT TTAGTGGCTT CACGCAGAAG TTGGCAGGAG CATGGGCTTC GGAGGCCTAT 120 AGCCCGCAGA TTNAAAGCCT GTGGTTTCCA CAGAAGCACC ACCTATCATA TTTGCCACAC 180 CAACTAAACT GACCTCCGAT TCCACAGTGT ATGATTATGC TGGGAAAAAC AAAGTTCCAG 240 25 AGCTACAAAA GTTTTTCCAG AAAGCTGATG GTGTGCCCGT CTACCTGAAA CGAGGCCTGC 300 CTGACCAAAT GCTTTACCGG ACCACCATGG CGCTGACTGT GGGAGGGACC ATCTACTGCC 360 30 TGATCGCCCT CTACATGGCT TCGCAGCCCA AAAACAAATG AGTTAGGCTG CAGAGGACTG 420 GTTTGTTTTT TGGCATAAAC CCTTTGAAGT TCCTTTTTCA TTGTTAAATT AAAATTTTTT TTTTTACTTG GATGGCTTAA CATTTTTGCA AGAAAAATAG GAAGATATGA AGATGATGTT 540 35 TTGGTTTGTT TATGAAATGC ATATGGCTTG TCAGAGCTCA TTCGACAGTT AAAGCCATTG 600 TTTAAAGAAA CGGTGCTTTG CTCTGTGTTT GTGCTCCTGA TTTCCCTGGA GGTTCTGGAT 660 40 GAAGGCTGAA CACAGGCTTG TTAATGTCAG TCTGTGCTGA GGACCTCAGG GACTTGAGGT 720 TGCATTTTTG AGCATGGGGT GCAGGAGCCT TTCTGGATTT GGATGTGGCT ATGGAAAGAA 780 CACAGAAGCC AAGGTCATGT GCATGAAATG AGGAGTTTGA GTTAGTCACC TCGGGGATTT 840 45 TTTCCATTTT GCAGTAAAAT GTTAAATTAA TGTAGCCTGC CTCTATTTGT TGGGCAGGTA 900 ATTTCAAAGG GTTATTTGCC TCATCTCCTA TCTTTAGTGA AATCTTATGT GTAATTGTGT 960 50 GTATTTATTC CACCGTGGGA ACAGAGAATA CCTGTTTAGT GTTGCACTTT AGACTGGTGT 1020 CTGTTTIGTT AATGCAGCTG TGCCACAAAT TCTCCTTTAT CTTTTAAAAA TGTTATAGCT 1080 TTAAATTTTG ATTTATTTTG ACTGTGGAAT AAATACATGA ATGAAAAATT TTAAGTTTGA 1140 55 AGTTCTTTGA ATGACCTTTC AGAGTAATTT CAGAACACCA GCAGCATCTT AAACCTGAGT 1200 CTAATTTCTT TCTTGTTAAT TAGGCACCAG ATAATCTTTA TAAAATGGTC TTAAAAGCTA 1260

GTAATAGGAG CTTAATGGCA ATKGATGATT ACCACAKGGT TTTTTATAAA AACCTGCCTG

1320

	CCCCTWAGTG	AAAGGTACCT	GTAACYCACA	GTYCATTTAG	ACACTAATTT	CCTYTGCYGT	1380
5	CATGATTGGK	AGACTTCACT	TACCCTATAT	TAATTTTGAA	AAAAGGTGGA	ATTTTATTAT	1440
_	ATATGAAGGA	ATAGTTTGTA	TCTTACCATA	GCACAGAACA	GTGACCTCTT	GCTCAGGATA	1500
	AGATGTGGTG	ATTTGAAAAT	ACTCATAGTA	GCCTTGCAGT	GATACCTCTC	TCNCTCTCTC	1560
10							
	(2) INFORM	ATION FOR SE	Q ID NO: 56	5:		,	
15 20	(i)	(B) TYP (C) STR	HARACTERIST GTH: 1507 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
20	(xi) SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 56:		
	GGAACGCAGA	GCGGAGCGTG	GAGAGCGGAG	CGAAGCTGGA	TAACAGGGGA	CCGATGATGT	60
25	GGCGACCATC	AGTTCTGCTG	CTTCTGTTGC	TACTGAGGCA	CGGGGCCCAG	GGGAAGCCAT	120
	CCCCAGACGC	AGGCCCTCAT	GGCCAGGGGA	GGGTGCACCA	GCGCCCCC	CTGAGCGACG	180
30	CTCCCCATGA	TGACGCCCAC	GGGAACTTCC	AGTACGACCA	TGAGGCTTTC	CTGGGACGG	240
	AAGTGGCCAA	GGAATTCGAC	CAACTCACCC	CAGAGGAAAG	CCAGGCCCGT	CTGGGGCGGA	300
	TCGTGGACCG	CATGGACCGC	GCGGGGGACG	GCGACGGCTG	GGTGTCGCTG	GCCGAGCTTC	360
35	GCGCGTGGAT	CGCGCACACG	CAGCAGCGGC	ACATACGGGA	CTCGGTGAGC	GCGGCCTGGG	420
	ACACGTACGA	CACGGACCGC	GACGGGCGTG	TGGGTTGGGA	GGAGCTGCGC	AACGCCACCT	480
40	ATGGCCACTA	CGCGCCCGGT	GAAGAATTTC	ATGACGTGGA	GGATGCAGAG	ACCTACAAAA	540
	AGATGCTGGC	TCGGGACGAG	CGGCGTTTCC	GGGTGGCCGA	CCAGGATGGG	GACTCGATGG	600
	CCACTCGAGA	GGAGCTGACA	GCCTTCCTGC	ACCCCGAGGA	GTTCCCTCAC	ATGCGGGACA	660
45	TCGTGATTGC	TGAAACCCTG	GAGGACCTGG	ACAGAAACAA	AGATGGCTAT	GTCCAGGTGG	720
	AGGAGTACAT	CGCGGATCTG	TACTCAGCCG	AGCCTGGGGA	GGAGGAGCCG	GCGTGGGTGC	780
50	AGACGGAGAG	GCAGCAGTTC	CGGGACTTCC	GGGATCTGAA	CAAGGATGGG	CACCTGGATG	840
	GGAGTGAGGT	GGGCCACTGG	GTGCTGCCCC	CTGCCCAGGA	CCAGCCCCTG	GTGGAAGCCA	900
	ACCACCTGCT	GCACGARAGC	GACACGGACA	AGGAYGGGCG	GCTGAGCAAA	GCGSAAATCC	960
55	TGGGTAATTG	GAACATGTTT	GTGGGCAGTC	AGGCCACCAA	CTATGGYGAG	GACCTGACCC	1026
	GGCACCACGA	TGAGCTGTGA	GCMCCGNGCA	CCTGCCACAG	CCTCAGAGGC	CCGCACAATG	1086
60	ACCGGAGGAG	GGGCCGCTGT	GGTCTGGCCC	CCTCCCTGTC	CAGGCCCCGC	AGGAGGCAGA	114

	TGCAGTCCCA GGCATCCTCC TKCCCCTGGG CTCTCAGGGA CCCCCTGGGT CGGCTTCTGT	1200
	CCCTGTCACA CCCCCAACCC CAGGGAGGGG CTGTCATAGT CCCAGAGGAT AAGCAATACC	1260
5	TATTTCTGAC TGAGTCTCCC AGCCCAGACC CAGGGACCCT NGGCCCCAAG CTCAGCTCTA	1320
	AGAACCGCCC CAACCCCTCC AGCTCCAAAT CTGAGCCTCC ACCACATAGA CTGAAACTCC	1380
10	CCTGGCCCCA GCCCTCTCCT GCCTGGCCTG GCCTGGGACA CCTCCTCTCT GCCAGGAGGC	1440
10	AATAAAAGCC AGCGCCGGGA AAAAAAAAAA AAAAAAAAA AAAAAAAA	1500
	AAAAAAN	1507
15		
20	(2) INFORMATION FOR SEQ ID NO: 57: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 450 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double	
25	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:	
20	GAATTCGGCA CGAGCAGTGT CCAACACTGT AGCTGGTGCC TGCCAGGTTC CCAGTGGCTG	60
30	GGGTCACCAG GTCTGAAGAG AGATGTGCTG GCTGCGGGCA TGGGSCCAGA TCYTCCTGCC	120
	AGTITICYTC TCCYTCTITC TCATCCAATT GCTTATCAGC TTCTCAGAGA ATGGTTTTAT	180
35	CCACAGCCCC AGGAACAATC AGAAACCAAG AGATGGGAAT RAAGAGGAAT GTGCTGTAAA	240
	GAAGAGTTGT CAATTGTGCA CAGAAGATAA GAAATATATG ATGAATAGAT AATTGAAAAG	300
40	AGATCCTCCA GAAAGAGCAG AAGGAAGTTT CTTCAATGGC TTCCTTCAGG ATTTTAATCA	360
40	TCCTTACAGC CTCTTTGAGA ATGATTGAAC TTCCAAATTC CCTGAAGTTA AAATTTTAAA	420
	TTCTATTAAA CATTTTTTCG AGTAAAAAA	450
45		
	(2) INFORMATION FOR SEQ ID NO: 58:	
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1147 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 58:	•
	GGCACGAGAC CCATTGAGCA GAAGGAGGCC AGGTGGGAAA GCTCCTGGGA AGAGCAGCCA	. 60
60	GACTGGACAC TGGGCTGCTT GAGTCCTGAG TCACAATTCA GAATTCCTGG GCTCCCTGGG	120

	TGCATTCTAT	CATTCCAGTT	GAAAGTTTGC	TTCCTTCCAG	TCATGTGGCT	CTTCATTCTA	180
	CTCTCCTTGG	CTCTCATTTC	AGATGCCATG	GTCATGGATG	AAAAGGTCAA	GAGAAGCTTT	240
5	GTGCTGGACA	CGGCTTCTGC	CATCTGCAAC	TACAATGCCC	ACTACAAGAA	TCACCCCAAA	300
	TACTGGTGCC	GAGGCTATTT	CCGTGACTAC	TGCAACATCA	TCCCCTTCTC	CCCTAACAGC	360
10	ACCAATCATG	TGGCCCTGAA	GGACACAGGG	AACCAGCTCA	TTGTCACTAT	GTCCTGCCTG	420
10	AACAAAGAAG	ACACGGGCTG	GTACTGGTGT	GGCATCCAGC	GGGACTTTGC	CAGGGATGAC	480
	ATGGATTITA	CAGAGCTGAT	TGTAACTGAC	GACAAAGGAA	CCTGGCCAAT	GACTTTGGTC	540
15	TGGGAAAGAC	TATCAGGCAC	AAAACCAGAA	GCTGCAAGGC	TCCCAAAGTT	GTCCGCAAGG	600
	CTGACCGCTC	CAGGACGTCC	ATTCTCATCA	TTTGCATACT	GATCACGGGT	TTGGGAATCA	660
20	TCTCTGTAAT	CAGTCATTTG	ACCAAAAGGA	GGAGAAGTCA	AAGGAATAGA	AGGGTAGGCA	720
15 20 25	ACACTTTGAA	GCCCTTCTCG	CGTGTCCTGA	CTCCAAAGGA	AATGGCTCCT	ACTGAACAGA	780
	TGTGACTGAA	GATTTTTTA	ATTTAGTTCA	TAAAGTGATG	CTACAACAGA	ATAATCACCA	840
25	TGACAACTGG	CCCCACACCT	CAGAGACTGA	TTCTGATCTC	CCAGGAATTC	TGAAGGTCCC	900
	TCTATCCTTG	ACAACAATCA	TTTGCAGCCA	GGTAGCAACG	GCAGTAGTCA	GAGGAGCTAT	960
30	GATAGACCAC	ACCCAAGCAA	GGCTGCCCTC	AAATAACATC	TCAAGATCTT	AGTTCTTATG	1020
50	CATTCCATCA	GTCAGAAGTG	AAGAAGAGGT	GGAGAATCTG	GATTGGGGAC	CAGGAAATCA	1080
	CTTGTATTTT	GTTAGCCAAT	AAATTCCTAG	CCAGTGTTGA	ATGAAAAAA	АААААААА	1140
35	AAAAAA						1147

40 (2) INFORMATION FOR SEQ ID NO: 59:

45

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 777 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 59:

50	GGCAGAGGCT	CCTCAGAAGG	GCGTGGGCTC	TCCAGTCTTC	CACAGTCCCC	ACCATGCCCT	60
	GTTGCCTTAC	CGCTGACGTA	GCTCACCCAT	CTTTTACTTG	CCTGGCTAAG	ATGCATGGCA	120
55	TYWCATTTCC	TCCTTGTTGC	ACTGCAGTCA	GTCCCTCACT	GCCCCATCT	CCTGGAAGAG	180
	GAGCATAAGC	TTTGCAAGGT	CAGCCACTTC	TCTGGGGTCA	CACTAGTTAC	ATCAAGACAG	240
	GACTCCAGCT	CATATGTGCC	AGTGCAGACA	CTCTTCATCC	ACCTGGGGCC	CTGGGCTTGG	300
60	GACCTGGYTC	CTTGCACAGC	AGARGACCCG	GAGGCTGAGA	GGAGCTTGCG	GTTGTGTCAT	360

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	AGTCACCTGG CCAGARGGAA CGTGAGCCCC TCCCAAGCTG CAGARGGARG GARCARGCGT	420
5	GGCTGTCAGC ACCGAGGTAG CAGAGAATTA ACATTCTTGT CAGCAGAGAA TGAAGCAGGA	480
3	ATATAATTAA AACTTTGCCC TTGGAATAGC TGATTCATTT GAATTTTATT CCACACGTTT	540
	GAAAGAGGAA AGAAAATGTG AAGACTTGCA GCCTGGTTCT CGCCTGGCCT GGGCTGGCCC	600
10	AGCTGTCAGG CCCGGTTCCT TTCTGAGCAT TCAGTCCACT GATGTTGACT GAGGGCCAGG	660
	AGAGACCCTC AGCAGGGTAT TACCATATCA GCCTCCTATC GCTGCTGGGA GAAATTACCA	720
15	TGAATTCAGT GGCTTAAAAC AACACACGAG CCTCTCTGAG CCTACCCTGG CTCAGGA	777
20	(2) INFORMATION FOR SEQ ID NO: 60: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1191 base pairs	
25	(B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 60:	
30	AAGANTGATT TTCCTTACTC TCCAAAGCGT CAGCATTTTG AAGTTTCTTT TATGAAAGTG	60
30 ° 3	GGGGCAAGAA TCAGGGTGAA AATGAGTGTA AACAAAGCCC ATCCTGTGGT CAGCACCCAC	120
	TGGAGGTGGC CAGCAGAGTG GCCTCAGATG TTCCTGCACC TGGCCCAGGA GCCCAGGACA	180
35	GAGGTCAAAT CTAGGCCCCT TGGTCTGGCT GGATTCATCA GGCAAGATTC GAAAACAAGA	240
	AAACCTCTAG AACAAGAAAC AATCATGTCT GCAGCAGATA CGGCACTGTG GCCCTATGGC	300
40	CATGGCAATC GTGAGCACCA AGAGAATGAG TTACAGAAAT ATCTCCAATA CAAAGACATG	360
10 15 20 25 30	CATCTCCTGG ACAGTGGACA GTCGCTGGGA CACACACACA CACTTCAAGG CTCACACAAC	420
	CTAACAGCCT TAAATATCTG AAGAAACAGA ATCACGACAT TAAGTCAGCA GAGGGAGAGG	480
45	TAGGCTGAAG CAGCAGGAGG CCAATTTTAT ATCCCACAGA TTTTTTTAAA AATGACTCCC	540
	CAGCAAGGGG TGGGGAGAAA GCCACTGATT TAGGAGAGTT CTTGGCTCAG CCAACCACTG	600
50	CGGTTATCTA CACGTTTTAC AAAGGCACRG AAGTAGAGAG GGGCTGCACT CACGACCCTC	660
- <u>-</u>	CCCAGGGCCC GCACAGCCAG ACACGGTGGG TTCTTCCTTT TTCCCTTCTG GCCTTGGTGG	72
	AATTCCTACC ACGGTGGCCT CTGCCTTTGG GACAATGCCT TCATGCTCAT CCCCGGGTCA	78
55	AGGATGGAGT CTGTTACCAT TTTCCAGGGG AAATTCCAAG GACCAGCCCC GCCTCATTAC	84
	GTTCACCCCA CAGGAAGGTG ATCTGGAAAG CCTGTAAACA CGTACTCTGG GTGGCTGAGT	90

GGTGTCACCA AGCTGCTTTT GTGCAGGGCT GAAGCACAGA CAAGAGGGCA GGCAGCTGCC

60

1200

	GGAGGCCTGA AGTGGGGAGA GATCCCCGCA GGCCTGCAGG AGCCAGGGAG AACCTCCAAC	1020
	TGGATCTAAA CTGTGGGACA GCCCAGGCGT GCCCCTCTTC ACATGGCTCC CAGGCTCCCT	1080
5	CAAAGCCCTT CCCAGGCCCT GCAGGAAGAG AGGGAGGGGTG AGGAGAGGCA GGGAGGGCAG	1140
	AGGTCGCCTG AAAGCCTGGG CTCCGAACTC CCTCAGCAGA GCTTTAAAGT G	1191
10		
10	(0) THE PROPERTY OF SEC. ID. NO. 44	•
	(2) INFORMATION FOR SEQ ID NO: 61:	
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1580 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 61:	
	CCCCGCCCCC CGCCCACGAA GGAAGTCGCT GCTGCTCCGG CGCGGACCCA GAGCCGGTTC	60
25	GGCGCGTCGA CTGCCCAGAG TCCGCGGCCG GGCGCGGGAG GAGCCAAGCC GCCATGGCCT	120
	ACCACAGCTT CCTGGTGGAG CCCATCAGCT GCCACGCCTG GAACAAGGAC CGCACCCAGA	180
	TIGCCATCTG CCCCAACAAC CATGAGGTGC ATATCTATGA AAAGAGCGGT GCCAAATGGA	240
30	CCAAGGTGCA CGAGCTCAAG GAGCACAACG GGCAGGTGAC AGGCATCGAC TGGGCCCCCG	300
	AGAGTAACCG TATTGTGACC TGCGGCACAG ACCGCAACGC CTACGTGTGG ACGCTGAAGG	360
35	GCCGCACATG GAAGCCCACG CTGGTCATCC TGCGGATCAA CCGGGCTGCC CGCTGCGTGC	420
	GCTGGGCCCC CAACGAGAAC AAGTTTGCTG TGGGCAGCGG CTCTCGTGTG ATCTCCATCT	480
	GTTATTTCGA GCAGGAGAAT GACTGGTGGG TTTGCAAGCA CATCAAGAAG CCCATCCGCT	540
40	CCACCGTCCT CAGCCTGGAC TGGCACCCCA ACAATGTGCT GCTGGCTGCC GGCTCCTGTG	600
	ACTTCAAGTG TCGGATCTTT TCAGCCTACA TCAAGGAGGT GGAGGAACGG CCGGCACCCA	660
45	CCCCGTGGGG CTCCAAGATG CCCTTTGGGG AACTGATGTT CGAATCCAGC AGTAGCTGCG	720
	GCTGGGTACA TGGCGTCTGT TTCTCAGCCA GCGGGAGCCG CGTGGCCTGG GTAAGCCACG	780
	ACAGCACCGT CTGCCTGGCT GATGCCGACA AGAAGATGGC CGTCGCGACT CTGGCCTCTG	840
50	AAACACTACC ACTGCTGGCG CTGACCTTCA TCACAGACAA CAGCCTGGTG GCAGCGGGCC	900
	ACGACTGCTT CCCGGTGCTG TTCACCTATG ACGCCGCCGC GGGGATGCTG AGCTTCGGCG	960
55 ·	GGCGGCTGGA CGTTCCTAAG CAGAGCTCGC AGCGTGGCTT GACGGCCCGC GAGCGCTTCC	1020
	AGAACCTGGA CAAGAAGGCG AGCTCCGAGG GTGGCACGGC TGCGGGCGCG GGCCTAGACT	1080
	CGCTGCACAA GAACAGCGTC AGCCAGATCT CGGTGCTCAG CGGCGGCAAG GCCAAGTGCT	1140

CGCAGTTCTG CACCACTGGC ATGGATGGCG GCATGAGTAT CTGGGATGTG AAGAGCTTCG

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960

60

	AGTCAGCCTT GAAGGACCTC AAGATCAAAT GACCTGTGAG GAATATGTTG CCTTCATCCT	1260
5	AGCTGCTGGG GAAGCGGGGA GAGGGTCAG GGAGGCTAAT GGTTGCTTTG CTGAATGTTT	1320
J	CTGGGGTACC AATACGAGTT CCCATAGGGG CTGCTCCCTC AAAAAAGGGAG GGGACAGATG	1380
	GGGAGCTTTT CTTACCTATT CAAGGAATAC GTGCCTTTTT CTTAAATGCT TTCATTTATT	1440
10	GAAAAAAAA AAAAATGCCC CCAAAGCACT ATGCTGGTCA TGAACTGCTT CAAAATGTGG	1500
	AGGTAATAAA ATGCAACTGT GTAAAAAAAAA AAAAAAAAAA	1560
15	AACTAGNCGG ACGCNTGGGT	1580
20	(2) INFORMATION FOR SEQ ID NO: 62: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1117 base pairs	
25	(B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 62:	
30	GGCACGAGGC GCGATGCAGC ACAGGCTAGA GGCTGCGCAA SGCGGGGGCC CGCCCC'IGGG	60
	ACCCTCCGGG CCGGGCGGTT TGGCCCCTTA GCGCCCGGGC GTCGGGGCGG TAAAAGGCCG	120
	GCAGAAGGGA GGCACTTGAG AAATGTCTTT CCTCCAGGAC CCAAGTTTCT TCACCATGGG	180
35	GATGTGGTCC ATTGGTGCAG GAGCCCTGGG GGCTGCTGCC TTGGCATTGC TGCTTGCCAA	240
	CACAGACGTG TTTCTGTCCA AGCCCCAGAA AGCGGCCCTG GAGTACCTGG AGGATATAGA	300
40	CCTGAAAACA CTGGAGAAGG AACCAAGGAC TTTCAAAGCA AAGGAGCTAT GGGAAAAAA	360
	TGGAGCTGTG ATTATGGCCG TGCGGAGGCC AGGCTGTTTC CTCTGTCGAG AGGAAGCTGC	420
	GGATCTGTCC TCCCTGAAAA GCATGTTGGA CCAGCTGGGC GTCCCCCTCT ATGCAGTGGT	480
45	AAAGGAGCAC ATCAGGACTG AAGTGAAGGA TYTCCAGCCT TATTYCAAAG GAGAAATCTT	540
	CCTGGATGAA AAGAAAAGT TCTATGGTCC ACAAAGGCGG AAGATGATGT TTATGGGATT	600
50	TATCCGTCTG GGAGTGTGGT ACAACTTCTT CCGAGCCTGG AACGGAGGCT TCTCTGGAAA	660
	CCTGGAAGGA GAAGGCTTCA TCCTTGGGGG AGTTTTCGTG GTGGGATCAG GAAAGCAGGG	720
•	CATTCTTCTT GAGCACCGAG AAAAAGAATT TGGAGACAAA GTAAACCTAC TTTCTGTTCT	780
55	GGAAGCTGCT AAGATGATCA AACCACAGAC TTTGGCCTCA GAGAAAAAAT GATTGTGTGA	840
	AACTGCCCAG CTCAGGGATA ACCAGGGACA TTCACCTGTG TTCATGGGAT GTATTGTTTC	900

CACTCGTGTC CCTAAGGAGT GAGAAACCCA TTTATACTCT ACTCTCAGTA TGGATTATTA

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	ATGTATTTTA ATATTCTGTT TAGGCCCACT AAGGCAAAAT AGCCCCAAAA CAAGACTGAC	1020
	AAAAATCTGA AAAACTAATG AGGATTATTA AGCTAAAACC TGGGAAATAG GAGGCTTWAA	1080
5	ATGACTGCCM GCTGGTGCRT GCTCACACTT GGCCCAC	1117
	•	
10	(2) INFORMATION FOR SEQ ID NO: 63:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 361 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:	
20	CCCACGCGTG CKGGCGCCTG GCAGCCACCG CCTGGGAGGT TACTGTAAGG CCCGCAGCTC	60
	CCGCCAGCTC CCGCGGACTS CTGCCGCCTC CTTACCATGA AGCCAGTAAG TCGTCGCACG	120
25	CTGGACTGGA TTTATTCAGT GTTGCTGCTT GCCATCGTTT TAATCTCCTG GGGCTGCATC	180
	ATCTATGCTT CGATGGTGTC TGCAAGACGA CAGCTAAGGA AGAAATACCC AGACAAAATC	240
	TTTGGGACGA ATGAAAATTT GTAACTCTTC TGGATTTAAT TATCTGAAAA TACAGTTCTT	300
30	TCCCTCATGC TTATGTAGAT ATAAAAATAA AATTCATAAT GCAAAAAAAA AAAAAAAAAA	360
	G	361
35		
	(2) INFORMATION FOR SEQ ID NO: 64:	
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1668 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 64:	
	GGCACGAGGT CTGCCAAGCT ATAGACCATG GCTGTGAACA CATTTGTGTG AACAGTGACG	60
50	ACTCATACAC GTGCGAGTGC TTGGAGGGAT TCCGGCTCGC TGAGGATGGG AAACGCTGCC	120
30	GAAGAAGGAT GTCTGCAAAT CAACCCACCA TGGCTGCGAA CACATTTGTG TTAATAATGG	180
	GAATTCCTAC ATCTGCAAAT GCTCAKAGGG ATTTGTTCTA GCTGAGGACG GAAGACGGTG	240
55	CAAGAAATGC ACTGAAGGCC CAATTGACCT GGTCTTTGTG ATCGATGGAT CCAAGAGTCT	300
	TGGAGAAGAG AATTTTGAGG TCGTGAAGCA GTTTGTCACT GGAATTATAG ATTCCTTGAC	360
60	AATTTCCCCC AAAGCCGCTC GAGTGGGGCT GCTCCAGTAT TCCACACAGG TCCACACAGA	420

	GTICACTCIG AGAAACIICA ACTCAGCCAA AGACATGAAA AAAGCCGTGG CCCACATGAA	480
	ATACATGGGA AAGGGCTCTA TGACTGGGCT GGCCCTGAAA CACATGTTTG AGAGAAGTTT	540
5	TACCCAAGGA GAAGGGGCCA GGCCCTTTCC ACAAGGGTGC CCAGAGCAGC CATTGTGTTC	600
	ACCGACGGAC GGGCTCAGGA TGACGTCTCC GAGTGGGCCA GTAAAGCCAA GGCCAATGGT	660
10	ATCACTATGT ATGCTGTTGG GGTAGGAAAA GCCATTGAGG AGGAACTACA AGAGATTGCC	720
10	TCTGAGCCCA CAAACAAGCA TCTCTTCTAT GCCGAAGACT TCAGCACAAT GGATGAGATA	780
	AGTGAAAAAC TCAAGAAAGG CATCTGTGAA GCTCTAGAAG ACTCCGATGG AAGACAGGAC	840
15	TCTCCAGCAG GGGAACTGCC AAAAACGGTC CAACAGCCAA CAGTGCAACA CAGATATCTG	900
	TTTGAAGAAG ACAATCTTTT ACGGTCTACA CAAAAGCTTT CCCATTCAAC AAAACCTTCA	960
20	GGAAGCCCTT TGGAAGAAAA ACACGATCAA TGCAAATGTG AAAACCTTAT AATGTTCCAG	1020
20	AACCTTGCAA ACGAAGAAGT AAGAAAATTA ACACAGCGCT TAGAAGAAAT GACACAGAGA	1080
	ATGGAAGCCC TGGAAAATCG CCTGAGATAC AGATGAAGAT TAGAAATCGC GACACATTTG	1140
25	TAGTCATTGT ATCACGGATT ACAATGAACG CAGTGCAGAG CCCCAAAGCT CAGGCTATTG	1200
	TTAAATCAAT AATGTTGTGA AGTAAAACAA TCAGTACTGA GAAACCTGGT TTGCCACAGA	1260
30	ACAAAGACAA GAAGTATACA CTAACTTGTA TAAATTTATC TAGGAAAAAA ATCCTTCAGA	1320
	ATTCTAAGAT GAATTTACCA GGTGAGAATG AATAAGCTAT GCAAGGTATT TTGTAATATA	1380
	CTGTGGACAC AACTTGCTTC TGCCTCATCC TGCCTTAGTG TGCAATCTCA TTTGACTATA	1440
35	CGATAAAGTT TGCACAGTCT TACTTCTGTA GAACACTGGC CATAGGAAAT GCTGTTTTTT	1500
	TGTAYTGGAC TTTACCTTGA TATATGTATA TGGATGTATG CATAAAATCA TAGGACATAT	1560
40	GTACTTGTGG AACAAGTTGG ATTTTTTATA CAATATTAAA ATTCACCACT TCAGAGRAAA	1620
	ΑΑΑΑΑΑΑΑ ΑΑΑΑΑΑΑΑΑ ΑΑΑΑΑΑΑΑΑ ΑΑΑΑΑΑΑΑΑ	1668
45	(2) INFORMATION FOR SEQ ID NO: 65:	
	(i) SEQUENCE CHARACTERISTICS:	
50	(A) LENGTH: 1353 base pairs(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 65:	
55	GGGTCGACCC ACGCGTCCGC CCACGCGTCC GGATGGCTGC GCTGTTGCTG AGACACGTTG	60
	GTCGTCATTG CCTCCGAGCC CACTTTAGCC CTCAGCTCTG TATCAGAAAT GCTGTTCCTT	12
60	TGGGAACCAC GGCCAAAGAA GAGATGGAGC GGTTCTGGAA TAAGAATATA GGTTCAAACC	180

المراكبة فللمشارك المستبيعة والمجوعين موادوا أأسترك أمكارك

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الهُمُّ الرياقة ويعان والمعالية المنطقة الالصيد في المنافقة من الما الما المنافقة المنافقة والما المنافظة المنطقة الم

	GTCCTCTGTC	TCCCCACATT	ACTATCTACA	GTTGGTCTCT	TCCCATGGCG	ATGTCCATCT	240
5	GCCACCGTGG	CACTGGTATT	GCTTTGAGTG	CAGGGGTCTC	TCTTTTTGGC	ATGTCGGCCC	300
-	TGTTACTCCC	TGGGAACTTT	GAGTCTTATT	TGGAACTTGT	GAAGTCCCTG	TGTCTGGGGC	360
	CAGCACTGAT	CCACACAGCT	AAGTTTGCAC	TTGTCTTCCC	TCTCATGTAT	CATACCTGGA	420
10	ATGGGATCCG	ACACTTGATG	TGGGACCTAG	GAAAAGGCCT	GAAGATTCCC	CAGCTATACC	480
	AGTCTGGAGT	GGTTGTCCTG	GTTCTTACTG	TGTTGTCCTC	TATGGGGCTG	GCAGCCATGT	540
15	GAAGAAAGGA	GGCTCCCAGC	ATCATCTTCC	TACACATTAT	TACATTCACC	CATCTTTCTG	600
15	TTTGTCATTC	TTATCTCCAG	CCTGGGAAAA	GTTCTCCTTA	TTTGTTTAGA	TCCTTTTGTA	660
	TTTTCAGATC	TCCTTGGAGC	AGTAGAGTAC	CTGGTAGACC	ATAATAGTGG	AAAAGGGTCT	720
20	AGTTTTCCCC	TTGTTTCTAA	AGATGAGGTG	GCTGCAAAAA	CTCCCCTTTT	TTGCCCACAG	780
	CTTGCCTACT	CTCGGCCTAG	AAGCAGTTAT	TCTCTCTCCA	TATTGGGCTT	TGATTTGTGC	840
25	TGAGGGTCAG	CTTTTGGCTC	CTTCTTCCTG	AGACAGTGGA	AACAATGCCA	GCTCTGTGGC	900
	TTCTGCCCTG	GGGATGGGCC	GGGTTGGGGG	GTGGGTTGGT	GAGGCTTTGG	GTGCCACTGC	960
	CTGTGGGTTG	CTGGCTTAAA	GGACAATTCT	CTTCATTGGT	GAGAGCCCAG	GCCATTAACA	1020
30	CCTACACAGT	GTTATTGAAA	GAAGAGAGGT	GGGGGTGGAG	GGGAATTAGT	CTGTCCCAGC	1080
	TAGAGGGAGA	TAAAGAGGC	TAGTTAGTTC	TTGGAGCAGC	TGCTTTTGAG	GAGAAAATAT	1140
35	ATAGCTTTGG	ACACGAGGAA	GATCTAGAAA	ATTATCATTG	AACATATTAA	TGGTTATTTC	1200
	TTTTTCTTGG	ATTTCCAGAA	AAGCCTCTTA	ATTTTATGCT	TTCTCATCGA	AGTAATGTAC	1260
	CCTTTTTTC	TGAAACTGAA	TTAAATACTC	ATTTTATCTT	TGAAAAAAA	AAAAAAAACC	1320
40	TNGGGGGGG	CCCCGGACCC	NAATTGGCCC	TAT			1353

45 (2) INFORMATION FOR SEQ ID NO: 66:

50

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1011 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 66:

55 CGGAAGAAG CAGCCATCCA GACATTTCAG AACACGTACC AGGTGTTAGC TGTGACCTTC 60

AATGACACAA GTGATCAGAT TATTTCTGGT GGAATAGACA ATGATATCAA GGTCTGGGAC 120

TGCGCCAGAA CAAGCTAACC TACACCATGA GAGGCCATGC AGATTCAGTG ACTGGCCTGA 180

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600

	GTTTAAGTTC TGAAGGCTCT TATCTTTTGT CCAATGCAAT GGACAATACA GTTCGTGTCT	240
	GGGATGTCCG GCCATTTGCC CCCAAAGAGA GATGTGTAAA GATATTTCAA GGAAATGTGC	300
5	ACAACTITGA AAAGAACCTT CTGAGATGTT CTTGGTCACC TGATGGAAGC AAAATAGCAG	360
	CTGGCTCAGC CGACAGGTTT GTTTATGTGT GGGATACCAC AAGCAGGAGA ATATTGTATA	420
10	AGCTGCCCGG CCATGCTGGC TCCATCAATG AAGTGGCTTT CCACCCTGAT GAGCCCATCA	480
10	TTATCTCAGC ATCGAGTGAC AAGAGACTGT ATATGGGAGA GATTCAGTGA AGATATGGAC	540
	TGGAAGACTC CAAGGCCGCT TGTCTTTGAG ACCTCAGACT GCATAAGTGA TGCCAAATGT	600
15	TGGATGTCCA GGYTAGCACC CTCCCTTCAG ATGACCATTG CTAGCAAGAA ACAGGAGGCG	660
	GTGGCCATAT TCCAAAAACC ACTTCTGTCC CATTICACCA GGATGACTAA GGCAAGCTCC	720
20	CTGTGGCCTC TAAAAACCAC CTGCCAGATT TCAGGGACTG TTTTTTTTTT	780
20	TITTCCTGTT TICTAATGCA GGCCCAATGT GACAAATTTG TTGGTTGGGA TITTTTTTT	840
_	TTTTTGTAAC TGGCTTGTAT GATATTTTCT TTCTGTATTT CTCTATATCA TTTTGTATTA	900
25	AAAGCCAAAT AGATGCCTIT TTACAAGARM AAAAAAAAAA AAAAAAAAA NNAAAAAAAA	960
	CTGGGAGGGG GGGCCCGGTA CCCAAATCGC CGGATATGAT CGTAAACAAT C	1011
30		
	(2) INFORMATION FOR SEQ ID NO: 67:	
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1193 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 67:	
	GCCCGGGGGG TGCGCACTGC GGGCGCATCC CTGCCCCGGC GCCGTCCGTG CCCGCGGGAC	60
45	CTGACAGCCG GGTCAGAGGG CGAACTGTGC TCAGGCCCGG GCTGGACGCA GAGCCAGAGC	120
49	TGTCCCCAGA GGAGCAGAGG GTCCTGGAAA GGAAGCTGAA AAAGGAACGG AAGAAAGAGG	180
	AGAGGCAGCG TCTGCGGGAG GCAGGCCTTG TGGCCCAGCA CCCGCCTGCC AGGCGCTCGG	240
50	GGGCCGAACT GGCCTGGGAC TACCTCTGCA GATGGGCCCA AAAGCACAAG AACTGGAGGT	300
	TTCAGAAGAC GAGGCAGACG TGGCTCCTGC TGCACATGTA TGACAGTGAC AAGGTTCCCG	360
55	ATGAGCACTT CTCCACCCTG CTGGCCTACC TGGAGGGGCT GCAGGGCCGG GCCCGAGAGC	420
33	TGACGGTGCA GAAGGCGGAA GCCTGATGCG GGAGCTGGAT GAGGAGGGCT CTGATCCCCC	480
	CCTGCCGGGG AGGGCCCAGC GCATCCGACA GNTGCTGCAG CTGCTCTCCT AGTGGGTTCA	540

GCGCGGGGC GGGCCGCTGC CCAGTGCAGG GCTGCCTCAG ACCACACAGG GTGCAGCTCC

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	TCCGGCGGTG	GGGGCCGGGT	TCACCAGCAG	GGCAGCGGCT	GAGCAAGGGC	TTTCAGCTCC	660
5	TCCGGTGGTG	GGGGCCGGGA	TCACCAGCAC	CAGAGCCTCG	CAAGGCCCC	TTCCCTCCTC	720
5	CAGACCCTCC	TTGGCCGGTG	ACGCTGTGAC	AGTGATGGCA	GGTTCAGTGC	CTTCAGCGCA	780
	GAGCGTGGAT	GCTCTGGAAT	CACCCGGACC	CCTGGCCTTG	GAGGGACCCT	CCAGCCCCAG	840
10	GAATCTGCTT	TGGAGGGAAA	TGTCTATTTT	TCTACCGGGA	ATATTTTAGA	GATTGGGGCA	900
	TGCTGGCTCC	TCCCGCCAGC	TGCAAACCTG	CACCTTCCGC	CTGATTCCCG	ATCCCCCTGC	960
15	GTGGGCCGCA	TTCCTGGTCC	CCTGCCTGCG	TCCATCGAGG	GGCCTGGCTG	TGGCCTGTTT	1020
13	TCCTTTGACC	CCACACAGCG	TCATTGCGGG	TCATGGGGAG	CCCCTGGTGG	GAGCTTGTGG	1080
	AGTCGGATCA	CGTACCTGTG	CAGAAACCGC	CTCTGTGGCT	GCATTTGAAA	TAAAACCCGA	1140
20	CCCAGCAGCA	ААААААААА	AAAAAANCNC	NAGGGGGGC	CCGGNACCCA	TTA	1193

25 (2) INFORMATION FOR SEQ ID NO: 68:

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(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 560 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 68:

GAATTCGGCA	CGAGTTGGCA	CATGATGCAA	AATGCATTTC	TCAGAGTAGA	TTGCAGTCAA	60
AAATGTTGGA	AACTACTAAG	CATGTGCARA	TAGCATGCAT	GCTGCTGCTG	ACCTGCCAGA	120
TATTTCTCCC	TTCCTCCCTT	TCTCCCTCAT	TTATTCATTC	ATTAACTGAT	TCATTCATCC	180
CATTAAAAAA	ATTATATGTA	TGTTTTGTGC	AAAGCACCCT	ACTCAAGGCT	GCGGGGTACA	240
AAAGTATATC	AGAAGCCTTG	GGCTTTGACM	WACTTCTCTG	TAGTAGTGCT	AGATTTGTGT	300
GGATCTGCCA	CACTTACTCC	AGGCCTCTTG	TGACCTGTGC	TTTGCATTAA	TCTCTTAGGC	360
TAAGCCACAT	ACCTTTTCAT	TATACAATCT	TTGCTGATGC	TAAGGACAGA	TTCCAAAGTG	420
CCCTCCTTAT	AATTTTTGTA	TTTAATGCAA	AGTGTAATCA	AGAATAGGCC	ATTGTTAGGT	480
CAATTGCTTT	TCTGTATTTA	TCTTTTCAAA	СААТАААТАА	TCAGTGGGAT	GAAAAAGGGC	540
CGGAAAAAA	ааааааааа					560

(2) INFORMATION FOR SEQ ID NO: 69:

60 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1657 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 69:

						•	
	CGGACNGAGC	CGCCGCCGGG	CACTTCCTGT	GGAGGCCGCA	GCGGGTGCGG	GCGCCGACGG	60
10	GCGAGAGCCA	GCGAGCGAGC	GAGCGAGCCG	AGCCGAGCCT	CCCGCCGTCG	CCATGGGCCA	120
	GAACGACCTG	ATGGGCACGG	CCGAGGACTT	CGCCGACCAG	TTCCTCCGTG	TCACAAAGCA	180
15	GTACCTGCCC	CACGTGGCGC	GCCTCTGTCT	GATCAGCACC	TTCCTGGAGG	ACGGCATCCG	240
15	TATGTGGTTC	CAGTGGAGCG	AGCAGCGCGA	CTACATCGAC	ACCACCTGGA	ACTGCGGCTA	300
	CCTGCTGGCC	TCGTCCTTCG	TCTTCCTCAA	CTTGCTGGGA	CANTGACTGG	CTGCGTCCTG	360
20	GTGTTGAGCA	GGAACTTCGT	GCAGTACGCC	TGCTTCGGGC	TCTTTGGAAT	CATAGCTCTG	420
	CAGACGATTG	CCTACAGCAT	TTTATGGGAC	TTGAAGTTTT	TGATGAGGAA	CCTGGCCCTG	480
25	GGAGGAGGCC	TGTTGCTGCT	CCTAGCAGAA	TCCCGTTCTG	AAGGGAAGAG	CATGTTTGCG	540
23	GGCGTCCCCA	CCATGCGTGA	GAGCTCCCC	AAACAGTACA	TGCAGCTCGG	AGGCAGGGTC	600
	TIGCTGGTTC	TGATGTTCAT	GACCCTCCTT	CACTTTGACG	CCAGCTTCTT	TTCTATTGTC	660
30	CAGAACATCG	TGGGGCACAG	CTCTGATGAT	TTTAGTGGCC	ATTGGTTTTA	AAACCAAGCT	720
	GGCTGCTTIG	ACTCTTGTTG	TGTGGCTCTT	TGCCATCAAC	GTATATTTCA	ACGCCTTCTG	780
35	GACCATTCCA	GTCTACAAGC	CCATGCATGA	CTTCCTGAAA	TACGACTTCT	TCCAGACCAT	840
55	GTCGGTGATT	GGGGGCTTGC	TCCTGGTGGT	GCCCTGGGC	CCTGGGGGTG	TCTCCATGGA	900
	TGAGAAGAAG	AAGGAGTGGT	AACAGTCACA	GATCCCTACC	TGCCTGGCTA	AGACCCGTGG	960
40	CCGTCAAGGA	CTGGTTCGGG	GTGGATTCAA	CAAAACTGCC	AGCTTTTATG	TATCCTCTTC	1020
	CCTTCCCCTC	CCTTGGTAAA	GGCACAGATG	TTTTGAGAAC	TTTATTTGCA	GAGACACCTG	1080
45	AGAATCAATG	GCTTCAGGAC	ATGGGTTCTC	TTCTCCTGTG	ATCATTCAAG	TGCTCACTGC	1140
	ATGAAGACTG	GCTTGTCTCA	GTGTTTCAAC	CTCACCAGGG	CTGTCTCTTG	GTCCACACCT	1200
	CGCTCCCTGT	TAGTGCCGTA	TGACAGCCCC	CATCAAATGA	CCTTGGCCAA	GTCACGGTTT	1260
50	CTCTGTGGTC	AAGGTTGGTT	GGCTGATTGG	TGGAAAGTAG	GGTGGACCAA	AGGAGGCCAC	1320
	GTGAGCAGTC	AGCACCAGTT	CTGCACCAGC	AGCGCCTCCG	TCCTAGTGGG	TGTTCCTGTT	1380
55	TCTCCTGGCC	CTGGGTGGGC	TAGGGCCTGA	TTCGGGAAGA	TGCCTTTGCA	GGGAGGGGAG	1440
55	GATAAGTGGG	ATCTACCAAT	TGATTCTGGC	AAAACAATTT	CTÄAGATTTT	TTTGCTTTAT	1500
	GTGGGAAACA	GATCTAAATC	TCATTTTATG	CTGTATTTTA	TATCTTAGTT	GTGTTTGAAA	1560
60	ACGTTTTGAT	TTTTGGAAAC	ACATCAAAAT	AAATAATGGC	GTTTGTTGTA	ааааааааа	1620

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	AAAAAAACTC GRGGGGGGC CCGGTACCCA AATCGCC							
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	(2) INFORMATION FOR SEQ ID NO: 70:							
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 711 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear							
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 70:							
	GGCACGAGCG AAGACCCTGT TCGGACCCTG CCCCGATTCC AGACTCAGGT AGATCGTCGG	60						
20	CATACCCTCT ACCGTGGACA CCAGGCAGCC CTGGGGCTGA TGGAGAGAGA TCAGGTATCC	120						
2 Ģ	CCCAGGGAGT AGGGGCTACC TTGAGGGGAT GATAGACCTC CCCCACTCCC AGTGKKACTC	180						
	TGGAAATATG AAGGAACTAG GGAGTGGAAG AGATTTCAGA GCTGGGGAGA GGAGTTCCTC	240						
25	CCTTCAAAGC CAGCAACTGC CTTTGGGGAA TGTCGGGGGG TCTCTCCTTT CTCCTGCTTG	300						
	TTTRAGGTGG TACACAGTCC CCCCTTCAMC TGGSGGGAAG CTGTNCCGGA CARACTCATC	360						
30	TCAGCTTTCC CTTGGGGCAG GATCGGGGC AGCAGCTCCA GCAGAAACAG CAGGATCTGG	420						
50	AGCAGGAAGG CCTCGAGGCC ACACAGGGGC TGCTGGCCGG CGAGTGGGCC CCACCCCTCT	480						
	GGRAGCTGGG CAGCCTCTTC CAGGCCTTCG TGAAGAGGGA GAGCCAGGCT TATGCGTAAG	540						
35	CTTCATAGCT TCTGCTGGCC TGGGGTGGAC CCAGGACCCC TGGGGCCTGG GTGCCCTGAG	600						
	TGGTGGTAAA GTGGAGCAAT CCCTTCACGC TCCTTGGCCA TGTTCTGAGC GGCCAGCTTG	660						
40	GCCTTTGCCT TAATAAATGT GCTTTATTTT CAAAAAAAAA AAAAAAAAAA	711						
45	(2) INFORMATION FOR SEQ ID NO: 71: (i) SEQUENCE CHARACTERISTICS:							
50	(A) LENGTH: 935 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear							
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 71:							
55	GGCACAGGGT GAAAGCCAGC TAAACCCCAA GTGGAGAAGT GAAAGACATG GTTGTTCCCA	60						
	TAAGTTTATT GCTCACATTA TGAAAGAAGC CATAGTCATG AGTGAACCAC TCCCTAGGTT	120						
	GATAAGGAAA CCAACACGGA AGATCTCTTT CTGGAAGAAG CAGCCAGCCT CGTGAAGGAG	180						
60	CGGCCCAGCC GCCGGGCCCG AGGGTCGCCT TTTGTTCGGA GTGGCACGAT TGTCCGTTCC	240						

	CAGACATTCT CGCCTGGAGC ACGAAGCCAG TATGTTTGCA GACTTTATCG TAGTGACAGC	300
5	GACAGTTCAA CGCTGCCCCG GAAGTCCCCC TTTGTCCGAA ATACTTTGGA AAGACGAACC	360
	CTTCGCTATA AGCAGTCATG CAGGTCTTCC CTGGCTGAGC TCATGGCCCG CACCTCCCTG	420
	GACTTGGAGC TGGATCTCCA GGCGTCGAGA ACACGGCAGA GGCAGCTGAA TGAGGAGCTC	480
10	TGCGCCCTCC GTGAGCTGCG GCAGCGGTTN GGAGGACGCC CAGCTCCGTG GCCAGACTGA	540
	CCTCCCACCC TGGGTGCTTC GGGACGAGCG GCTCCGTGGC CTGCTGCGGG AGCCGAGCGG	600
15	CAGACAAGAC AGACCAAACT TGACTACCGT CATGAGCAGG CGGCTGAGAA GATGCTGAAG	660
	AAGGCCTCCA AGGAGATCTA CCAGCTGCGT GGCAGAGCCA CAAAGAGCCC ATCCAAGTGC	720
	AGACCTTTAG GGAGAAGATA GCATTCTTCA CAAGGCCAAG GATCAACATA CCTCCTCTCC	780
20	CAGCCGACGA CGTCTGATGG AGTGCATTGT GCACATGAAG TATTTATCCA CCTGTTTTAT	840
	TTTCATGAAG TTCTTAGACT AGCTGAATTT GTCTTTAAAA TATTTGTGCA AAGCTATTAA	900
25	TATACACATT TTGTAAAAAA AAAAAAAAA AAACT	935
	(2) INFORMATION FOR SEQ ID NO: 72:	
30	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 504 base pairs (B) TYPE: nucleic acid	
35	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 72:	
40	GCAGGGGCGA GGGGYTGGGG ACCGCGGGGC GGACGGGAGC GAGTATGTCC GCTCTGACTC	60
70	GGCTGGCGTC TTTCGCTCGC GTTGGAGGCC GCCTTTTCAG AAGCGGCTGC GCACGGACTG	120
	CTGGAGATGG TGGAGTCCGT CATGCCGGTG GTGGTGTGCA CATTGAGCCC CGGTATAGAC	180
45	AGTTCCCCCA GCTGACCAGA TCCCAGGTGT TCCAGAGCGA GTTCTTCAGC GGACTCATGT	240
	GGTTCTGGAT TCTCTGGCGC TTTTGGCATG ACTCAGAAGA GGTGCTGGGT CACTTTCCGT	300
50	ATCCTGATCC TTCCCAGTGG ACAGATGAAG AATTAGGTAT CCCTCCTGAT GATGAAGACT	360
50	GAAGGTGTAG ACTCAGCCTC ACTCTGTACA AGAGCCAGGT GAGAATTTCA AGGATTATCG	420
	ACTTCATATT GCACATTAAA GTTACAAATT AAAGTGGCTT GGTCAAGAAT GARAAAAAAA	480
55	AAAAAAAATT GGGGGGGGC CCCN	504

5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 620 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 73:	
10	GAATTCGGCA CGAGGAGGAG GGGAGGCGGG GTAAGTTTGG TGGGAAACTC TGTAATTTCC	60
	WITTTACTT TCACAGCAAT AGTGCAGAAT CCAGAATGGA TGTCCTCTTT GTAGCCATCT	120
15	TTGCTGTGCC ACTTATCCTG GGACAAGAAT ATGAGGATGA AGAAAGACTG GGAGAGGATG	180
	AATATTATCA GGTGGTCTAT TATTATACAG TCACCCCCAG TTATGATGAC TTTAGTGCAG	240
	ATTICACCAT TGATTACTCC ATATTIGAGT CAGAGGACAG GCTGAACAGG TTGGATAAGG	300
20	ACATAACAGA AGCAATAGAG ACTACCATTA GTCTTGAAAC AGCACGTGCA GACCATCCGA	360
	AGCCTGTAAC TGTGAAACCA GTAACAACGG AACCTCAGAG TCCAGATCTG AACGATGCCG	420
25	TGTCCAGTTT GCGAAGTCCT ATTCCCCTCC TCCTGTCGTG TGCCTTTGTT CAGGTGGGGA	480
23	TGTATTTCAT GTAGAAGGTG GAAGAAGGCT GCTATGACTC TTTGGATGGG AGTCTGGCAA	540
	GAGGAAATTG GAAGATAAAA TAAATAATAA GTGAAATAAA AAAAAAAA	600
30	GGGGGGCCC GGTACCCAAT	620
35	(2) INFORMATION FOR SEQ ID NO: 74:	
40	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 581 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 74:	
45	ACAAGGTGTG TGTAAAGTTT ATGTTTGTAA ACTGAATTCT ATCTTAAATC CAAAAAGAAC	6
	TCGGGAGTAA TTCATTTTG TAGCATAAAG ATCCCTAAGT TTTATTTTGA AATATCTGAT	12
50	TTTTACACGT TAAAAAATAA CAGGGCATCG AGAGGATTCC TAGGTGACAT CCAGACTCCT	18
50	TTAGCTTTGT GTGTGTGGCA CCGGTTAGTC TGCTTCTCTC TCCTTTCTTG CACTGCTTCA	24
	CACAGCCATG CCCTGCCAGC CCGGGCAGGT GCCTTCCTGT CAATGTACAT TTGGGCTTCT	30
55	GCTCATGCTG CCCTCCCTCC CCTCCCCTGC CTCCCAACCC CGCCCCTTTT GTTCCTCCAT	36
	GGAGTACTTC CATGGGTGTG CCTCCCCCAG CCAAGCCATA ATAGGTGGTT TCCCCTTCGC	42

TTCTGTAGCC CTTGCAGACA TCCTCTGTTT ACAGTAGGTG TTGACTTACT TCCCCTCTCC

CCGSTAAAGC	CATAAACTCC	TTAAGGACAG	GTAGCATTCT	TAGTATCTTC	GTTCTTCTCA	540
ATGACCAGTA	GACCATTAAA	CATGTAGCAA	ACAAATGTGA	A		581

(2) INFORMATION FOR SEQ ID NO: 75:

10 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1843 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 75:

	AAACCCAACN	CCCTCCGGTC	CCCNAAAGAA	AGCCCAGCCC	AAATCCCAAG	CCGGCAGTGA	60
20	GCCCGCGAAC	AAGGCCCTCA	AGACGCCCAG	NCGAACAAGC	AGCCCCCAGG	AGGCCCCGCA	120
	AGAGAACTCC	CTGGCGGCCC	AAGCGGGCAG	CTTCTGTGCG	GCAGAACTCA	GCCACCGAGA	180
25	GCGCAGACAG	CATCGAGATT	TATGTCCCGG	AGNCCCAGAC	CAGGCTCTGA	GACCATGCAG	240
23	GAGGAAAGAA	ACGATTTTAA	ATCATTAAAA	ACACAAAAAC	TAAGTGCGAA	CGGAACAGAG	300
	TTTTCTCAAC	CTTTGCTATG	GTTATTCTGT	CTAGAGACCC	TGAGCCAACT	TTCAAATTGA	360
30	CGCATACAAG	GGCTCACAAT	TIGGCTTTTT	TGGGTCCCTC	CCAGCTTTAG	GTTATGAAGA	420
	TTTTACTCAC	AAAAAAAATC	AACAAAAATC	ACGAAACTAG	AAAACTTTTT	TTTTCCTCTT	480
35	GCTGGCCGTG	GTGGACTAGA	TAGATGGACG	TCGGCAACTC	CCGGCCCAGC	CTCCATACTG	540
33	CGGTCTTTTT	ACTCGTTCTA	TCTGATGAGA	ACTCACACTA	GCTTGTTTAC	AAGATGACGA	600
	CAGTCCAAGG	GCAGCCTTGG	GCACCTGCCA	TGTCCCTCCT	TTCCCCAGCT	ATCCCCGCTC	660
40	TGACCTTGAT	TTTCATTCTT	ATGTTTTCT	CTTTTCCCTT	CAGAGCTCAC	ACAGTGGTCA	720
	CCATTGTGGC	AAGCGGCTTT	CTGGGTCTCA	GCCCTCTCTG	CGGTTGAGGG	CCCAGAGGAC	780
15	AGAGAGATGG	ACATGCGTCC	CCTCCCTCCC	CCCGCCAAGT	GCTCACACAC	AACCTCACGC	840
45	GCACACACAC	ACACGCAGAT	GGAGGCGCCT	' CACTGGGAGG	TGCCCGCCA	GCCCTGGGCA	900
	GTGTCAGGCA	GGACTCACTC	ACCGCTGAGC	AGATGAGAGA	AGTTTTAGTC	TIGGCGGGTG	960
50	GAAATGAGAC	GAAGCCACAG	TTATCACACT	CCAGACTCCT	GCCCTTTTAT	TTTCTCCAGC	1020
	CCCTTCTTCC	TTCAGCAAAA	TCTAGGACTC	CCGAGTGGCT	TCCAGGGGG	CGTCAGTCCT	1080
<i></i>	CAGCCGCGCC	TGTGTCCGGT	GCCCGAGGGG	ceeeceece	G TGTCTGTATC	TATGTGTACA	1140
55	TATGCACATA	GACCTTAGAG	; TGTATAGTTA	ACAAACGCCC	ATCTGCTCAC	CCATGCCCAC	1200
	CCAGCGCCGC	CGCCGCTGGC	TCTCGGGGC	A CCTGGCAGG	A GGCGGGTGTC	G TGAATAGCAT	1260
60	ATATTTTAC	ATGTACTATA	TCTAGGTGTG	G TGTACAAGTO	G TGTGTAAAA	TATATACCTT	1320

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	GTGTGTAAGC AGCCCTTTTT TTTTTTGGTC TCCACCCCCC TCCCCCGCC CCGCACTCCT	1380
5	AAGGGCCCAT CTGCCCAGCC TCTGAGTTTT CTGTTCTATT TTTTTTTTAA CCCCAATTAT	1440
,	CCTTCTCTCT CTCCTGCCCC CGCATCCCAC TCCCAGGGTG TCACGAGCCC TGAGCTGCAA	1500
	TGGCCCGGGC CTGCAGGGCG GGGTAGGGGA GGGCARGGCT SAGCCCCGAA GCCAGCTCAG	1560
0	TACCTGAGGG GCTGCTCTAT GCTGTGTATG CGCCTCTCTG GCATCCGAGA CATCCTCTTG	1620
	GTGGCGCTTG CTNGCAGGGG ACCCCCCCC CGTCCCCAGG TGAACCAAGG GTCTGCTCCG	1680
15	GGGCCCATTT CCAGCTTGGC CGCCGTCTGT GACCTTGGGC AAGTCACTTG ACCTCTGTGT	1740
13	GCCTCAACTT CCTCCTCTGT AAAACGCGGA CAGTCCCTGC CCCTCCCTAC CTCACAGGCA	1800
	TGTTGTGAGA ATAAATGAGG TAACGTGTAA AAAAAAAAAA	1843
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	(2) INFORMATION FOR SEQ ID NO: 76:	
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1441 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
30	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 76:	
	TCGACCCACG CGTCCGGCTC CCCGAGCCCT GCCAACCATG GTGAACTTGG GTCTGTCCCG	60
35	GGTGGACGAC GCCGTGGCTG CCAAGCACCC GGGACTCGGG GAGTATGCCG CATGCCAGTC	120
	ACACGCCTTC ATGAAGGGCG TTTTCACCTT CGTCACAGGC ACCGGCATGG CCTTTGGCTT	180
40	GCAGATGTTC ATTCAGAGGA AGTTTCCATA CCCTTTGCAG TGGAGCCTCC TAGTGGCCGT	240
	GGTTGCAGGC TCTGTGGTCA GCTACGGGGT GACGAGAGTG GAGTCGGAGA AATGCAACAA	300
	CCTCTGGCTC TTCCTGGAGA CCGGGCAGCT CCCCAAAGAC AGGAGCACAG ATCAGAGAAG	360
45	CTAGGAGAGC TCCAGCAGGG GCACAGAGGA TTGGGGGCAG GAGGAGTCTG GAACACAGCC	420
	TTCATGCCCC CTGACCCCAG GCCGACCCTC CCCACACCCT AGGGTACCCC AGTCGTATCC	480
50	TCTGTCCGCA TGTKTGGCCA GGCCTGACAA ACACCTGCAG ATGGCTGCTG CCCCAACCTG	540
	GGACCTGCCC AGRAGGTTGG AGCAGAAAGG GCTCTCCCTG GGGTGGTGTT TCTCCTCTAG	600
	GGTATTGGGA TGCATGTTCT GCACTGCCAG CAGAGAGGGT GTGTCTGGGG GCCACCACCT	660
55	ATGGGACACG GGGTCGAAGG GGCCTGTACA CTCTGTCATT TCCTTTCTAG CCCCTGCATC	720
	TCCAACAAGT CCAAGGTGAC AGCTGGTGCT AGGGGCGTGG GGTTAATAAA TGGCTTATCC	780
	TTCTCTCCAC CCAAGTTTCC ACCTGACCAG GTGAAAAACA AATCAGAAGG GTAAGATGAT	840

	GACAGGTCAC	ATGAAACCTT	TATTACCCTA	CAGTTGATAT	ATGAGGATCA	CATGCAAGTT	900
	ACATACTGAG	GATGTACAGG	GAAGTTCCCA	GCGCTGAACC	CCAGAATTAG	ACGTTCGCAT	960
5	CAGCCCCGTA	GGCCACGTGG	ACACCACCAC	AGCCTCTCTG	TATGGGGGTC	TGCCTCTGTA	1020
	GCACTTGGCA	TGTAGGGGCA	GAGCAAAAGG	GGCCANGCTG	GCCAGAGCCT	GGCTGCTGGG	1080
10	NAGARGAGGG	ACTTGTGGGS	CACGCCACNT	GCCTATCATT	CCCCAYTCAT	CTATTAGCCA	1140
	AAGTCACTCC	CCAGAGGCAG	AGCTAGCCCG	TTGTAGCCGT	GTCTGTGTGG	AGGGAAAGCT	1200
	TCTGAGTGGG	CAAGCCTACA	CACAGCCCCG	AGCCCCAAGA	GGAGGAAGAG	GTĠGAGACCA	1260
15	GACGGAACCT	CCACAAGTCC	ATCATGGTTA	CAGCTGGCTT	CCCCGCAGCA	CCGAAGACCC	1320
	ACAGCATNGG	CCCTGCTGCC	CCCGACCCAG	CTCAGCTGCC	ANGCCTCACC	TTGCCAGGAA	1380
20	TTGAAAGAAA	GTTATTGAGT	ACTAATTGGC	CTCAGAGTNA	CAGGAAGCTC	AAGTTAAAGT	1440
	G						1441
25							

(2) INFORMATION FOR SEQ ID NO: 77:

(i) SEQUENCE CHARACTERISTICS:

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(A) LENGTH: 910 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 77:

	GGCAGAGCTG	GCCTTCGACT	CGCTATGTCC	ACTAACAATA	TGTCGGACCC	ACGGAGGCCG	60
	AACAAAGTGC	TGAGGTGAGG	ACCCCAGCGT	CGTGGGCACG	GGTTCGGGTT	GTGGGTGTGG	120
40	ATCGGGGCCC	TGGGAAGCGC	CTGTCTATCC	CGGGGGCAGG	ACCTGAGCGC	CCCTGACCCT	180
	CGAGCCTGTC	GCAGGTACAA	eccccecce	AGCGAATGTA	ACCCGGCCTT	GGACGACCCG	240
45	ACGCCGGACT	ACATGAACCT	GCTGGGCATG	ATCTTCAGCA	TGTGCGGCCT	CATGCTTAAG	300
	CTGAAGTGGT	GTGCTTGGGT	CGCTGTCTAC	TGCTCCTTCA	TCAGCTTTGC	CAACTCTCGG	360
	AGCTCGGAGG	ACACGAAGCA	AATGATGAGT	AGCTTCATGT	GAGACTTGCC	CTACAGAACA	420
50	AGTGACTCTT	GAGTAAGGGG	TGGGGGGACC	CCAGCCTGGC	CATCCTAGAC	TGACACCTCT	480
	CTCCTGTCTT	CATGCTGTCC	ATCTCTGCCG	TGGTGATGTC	CTATCTGCAG	AATCCTCAGC	540
55 [.]	CCATGACGCC	CCCATGGTGA	TACCAGCCTA	GAAGGGTCAC	ATTTTGGACC	CTGTCTATCC	600
55	ACTAGGCCTG	GGCTTTGGCT	GCTAAACCTG	CTGCCTTCAG	CTGCCATCCT	GGACTTCCCT	660
	GAATGAGGCC	GTCTCGGTGC	CCCCAGCTGG	ATAGAGGGAA	CCTGGCCCTT	TCCTAGGGAA	720
60	CACCCTAGGC	TTACCCCTCC	TGCCTCCCTT	CCCCTGCCTG	CTGCTGGGGG	AGATGCTGTC	780

	CATGTTTCTA GGGGTATTCA TTTGCTTTCT CGTTGAAACC TGTTGTTAAT AAAGTTTTTC	840
5	ACTCTGAAAA AAAAAAAAA AAAAAAAAAC TYGRGGGGGG GCCCGGAACC CAATTCSCCG	900
	GATAGTGAGT	910
0	(2) INFORMATION FOR SEQ ID NO: 78:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2776 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 78:	
-0	TCGACCCACG CGTCCGGGCG GGCAGTGATG GCGGCTGGTG ATGGGGACGT GAAGCTAGGC	60
	ACCCTGGGGA GTGGCAGCGA GAGCAGCAAC GACGGCGGCA GCGAGAGTCC AGGCGACGCG	120
25	GGAGCGGCAG CGRAAGGGGG AGGCTGGGCG GCGGCGGCGT TGGCGCTTCT GACGGGGGGC	180
	GGGGAAATGC TGCTGAACGT GGCGCTGGTG GCTCTGGTGC TGCTGGGGGC CTACCGGCTG	240
30	TGGGTGCGCT GGGGGGGGG GGGTCTGGGG GCCGGGGGCCG GGGCGGCGA GGAGAGCCCC	300
	GCCACCTCTC TGCCTCGCAT GAAGAAGCGG GACTTCAGCT TGGAGCAGCT GCGCCAGTAC	360
	GACGGCTCCC GCAACCCGCG CATCCTGCTC GCGGTCAATG GGAAAGTCTT CGACGTGACC	420
35	AAAGGCAGCA AGTTCTACGG CCCGGCGGGT CCATATGGAA TATTTGCTGG TAGGGATGCC	480
	TCCAGAGGAC TGGCCACATT TTGCCTAGAT AAAGATGCAC TTAGAGATGA ATATGATGAT	540
10	CTCTCAGATT TGAATGCAGT ACAAATGGAG AGTGTTCGAG AATGGGAAAT GCAGTTTAAA	600
	GAAAAATATG ATTATGTAGG CAGACTCCTA AAACCAGGAG AAGAACCATC AGAATATACA	660
	GATGAAGAAG ATACCAAGGA TCACAATAAA CAGGATTGAA CTTTGTAAAC AACCAAAGTC	720
15	AGGGGCCTTC AGAACTGCAA TTCTTACTCC CTTTCACAGA CTGTCCGGAG TCTTTGGGTT	780
	TGATTCACCT GCTGCGAAAA ACATTCAACA AATTGTGTAC AAGATAAATT AATCTCACTA	840
50	TGAAGATTTG AATAACTAGA CATTATTTAT GCTGCCAAAC TCATTTGTTG CAGTTGTTTG	900
	TAATGTCTAG TGGGGCTTCA TCATCCTGAA AAGAAGGAGA CAGGGATTTT TTTAAAGAGC	960
	AAGAAAGTCA CAATATTACT TCTTTCCTTC CTTTTTTCCT TCTTTCCTTT CTTCTT	1020
55	TTTCTTTCTT TTTAAAATAT ATTGAAGACA ACCAGATATG TATTTGCTAC TCAAGTGTAC	1080
	AGATCTCCTC AAGAAACATC AAGGGACTCC TGTGTCACAT ACTGTGTTTT TATTTTAACA	1140
.	TGGGTGAGGG AGGCGACCTG ATCAGGGGAG GTGGGGGTAC ACATCAATTT GAGTTGTTCA	1200

	GGCTACTGAA ACATTAAAAT GTGAATTCCC AAACTTTTCT TTTTGGCTTT GTCAGGGAAA	1260
	AGAAAAATAT CTTTATAAAG AAATCTTTGG AAATTAGGAG AAGGAATTTC AGGTGGGTTT	1320
5	AAGTCAGAGC TAGTTCCCCA ACAGAAAGAT CATTTGAAAC CAGTTTTTAT CCCTTCTCTT	1380
	TCCTTCCCTT TCCCTAAATC AAATCAATAT TAATTGTGCC TTATTTCACT TAACATAGAC	1440
10	TTGAATTATT TTTAGGGAAA GCCCCTATAA TGAATTCAGA AATCACTACA AGCAGCATTA	1500
10	AGACTGAAGT TGGAATATTC TGTTGACCAT AAAACCTTGA TATCATTCTG TGTATATAGA	1560
	ATGTAAAAGG AATATTACAG TGTTAACTGC CATATATGTA ATATACACAA ACTCAATTAG	1620
15	CATTGTAATG GCCAAATGCA TTCCCCCATG CTTTTCTGTT TTCAAAAAAA TTGAAAAACA	1680
	AATCAACTCT TATCCCCAAC AGCTGCCTAA TTTTAGGAGT CTGACCCTCC ACATCTCACT	1740
20	CGTGTGGGTG CATGGGGCTG TGGAGTGGGT GTCAGTATGG ATGTGTCTGA ATGTGTGAGG	1800
20	CCTTGGAAGG GACTCTTTCT GCAGATACTG TAAATACAAG TACCATTTTA ATAAAGCATG	1860
	TACAATAAAC CAAAATAAGC TTGAGTTGGA CTTTATATAC AGAACTGTAA GCCAGTGCAT	1920
25	TATGATACAG TTGTAAGATT GTGCATTTGA TTCAAGATAA GGAAAAATCT TGGAAATGAA	1980
	AAGCAGGCAC KGGTTAACCA AGTTGTACAC ATTGTACCAC ATTCAGCATA ACTTTAGGAA	2040
30	GAAATTCCAC TTTGTGAACA TTCTCCAGAA ATCCAAGATT ATTCAGGTAA GAATTGGTAT	2100
30	ATTAAATGTA CATCTTTTTA CTTTCTATTT TGATGCCAAC TGATTATACT AGACAATTAG	2160
	CACTCCAGGT GGTTATTGAA CACAAAACAG TAAAAGAATA TTGCACTGAT AGATACTAAA	2220
35	TTATTATTTT ATTAGGTTGA AAAAGCCCTT ACTAAAAGCC CCTCATATAT CAATTACTTT	2280
	ATTTCATTAT GACTACTTAG GTTCCGGGCT GGGGACAAGT TCACTTAAAA AGGCAATGTT	2340
40	ATTTAACAGG TCACCAGTTA AGACTTCTGC TTTGTAGATA CATGCAGAAG CCATCAAACA	2400
10	AGGGGGRGCT TTTAACTGCA ACAATAAGCT AAAGTATGTA AAATACTACA TTCTATTCAG	2460
	TCTTGGAGTG TTTTGTAGAA AGTTATCTTC AGCCAAATCT TTGCTGAAGA CTGGTTGTGG	2520
45	AGTGTTGGTA AATGCTTTGT GTTTTTATGT AAAATATTTT CTAAACAAAA AATGTTAAAA	2580
	GTACATGTCC TCTGTAGTAA ACTGATATCT ATATATATGA ATCATTCAAG CCTAAAGTCT	2640
50	AGTAATAAAC TGTACTTGTG AATAGAGAAA CCCTAAATAT TCATGCAGWA AAAATTATGC	2700
50	GGTCTGTTAA GAAAAATGAG TAATTTGTGT TTTGGACTTG AAATAAACAG TGTTCTGTAG	2760
	ATAATICCTC AACTIC	2776

60 (i) SEQUENCE CHARACTERISTICS:

⁽²⁾ INFORMATION FOR SEQ ID NO: 79:

(A) LENGTH: 1525 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 79:

		_		_	**	•	
	CCGCTGCTGA	TAACTATGGC	ATCCCCCGGG	CCTGCAGGAA	TTCGGCACGG	AGCTACGGCG	60
10	CCGCCTGGCT	CCTGCTGNCA	CCTGCAGGCT	CGTCGCGGGT	GGAGCCCACC	CAAGACATCA	120
	GCATCAGCGA	CCAGCTGGGG	GGCCAGGACG	TGCCCGTGTT	CCGGAACCTG	TCCCTGCTGG	180
15	TGGTGGGTGT	CGGCGCCGTG	TTCTCACTGC	TATTCCACCT	GGGCACCCGG	GAGAGGCGCC	240
13	GGCCGCATGC	GGASGAGCCA	GGCGAGCACA	CCCCCTGTT	GCCCCTGCC	ACGCCCAGC	300
	CCCTGCTGCT	CTGGAAGCAC	TGGCTCCGGG	AGCSGGCTTT	CTACCAGGTG	GGCATACTGT	360
20	ACATGACCAC	CAGGCTCATC	GTGAACCTGT	CCCAGACCTA	CATGGCCATG	TACCTCACCT	420
	ACTCGCTCCA	CCTGCCCAAG	AAGTTCATCG	CGACCATTCC	CCTGGTGATG	TACCTCAGCG	480
25	GCTTCTTGTC	CTCCTTCCTC	ATGAAGCCCA	TCAACAAGTG	CATTGGGAGG	AACATGACCT	540
	ACTTCTCAGG	CCTCCTGGTG	ATCCTGGCCT	TTGCCGCCTG	GGTGGCGCTG	GCGGAGGGAC	600
	TGGGTGTGGC	CGTGTACGCA	GCGGCTGTGC	TGCTGGGTGC	TGGCTGTGCC	ACCATCCTCG	660
30	TCACCTCGCT	GGCCATGACG	GCCGACCTCA	TCGGTCCCCA	CACGAACAGC	GGACTKTCGT	720
	GTACGGCTCC	ATGAGCTTCT	TGGATAAGGT	GGCCAATGGG	CTGGCAGTCA	TGGCCATCCA	780
35	GAGCCTGCAC	CCTTGCCCCT	CAGAGCTCTG	CTGCAGGGCC	TGCGTGAGCT	TTTACCACTG	840
	GGCGATGGTG	GCTGTGACGG	GCGCCGTGGG	CGTGGCCGCT	GCCCTGTGTC	TCTGTAGCCT	900
	CCTGCTGTGG	CCGACCCGCC	TGCGACGCTG	GGACCGTGAT	GCCCGGCCCT	GACTCCTGAC	960
40	AGCCTCCTGC	ACCTGTGCAA	GGGAACTGTG	GGGACGCACG	AGGATGCCCC	CCARGGCCTT	1020
	GGGGAAAAGC	CCCCACTGCC	CCTCACTCTT	CTCTGGACCC	CCACCCTCCA	TCCTCACCCA	1080
45	GCTCCCGGG	GTGGGGTCGG	GTGAGGGCAG	CAGGGATGCC	CGCCAGGGAC	TTGCAAGGAC	1140
	CCCCTGGGTT	TTGAGGGTGT	CCCATTCTCA	ACTCTAATCC	ATCCCAGCCC	TCTGGAGGAT	1200
	TTGGGGTGCC	CCTCTCGGCA	GGGAACAGGA	AGTAGGAATC	CCAGAAGGGT	CTGGGGGAAC	1260
50	CCTAACCCTG	AGCTCAGTCC	AGTTCACCCC	TCACCTCCAG	CCTGGGGGTC	TCCAGACACT	1320
	GCCAGGGCCC	CCTCAGGACG	GCTGGAGCCT	' GGAGGAGACA	GCCACGGGG1	CGTGGGCTGG	1380
55	GCCTGGACCC	CACCGTGGTG	GGCAGCAGGG	CTGCCCGGCA	GCTTGGTG	ACTCTGCTGG	1440
	CAGCAAATAA	AGAGATGACG	GCAAAAAAA	AAAAAAAA	AAAAAAAA	AAAAAAAA A	1500
	ААААААА	AAACCCACCG	TCCGC				1525

(2) INFORMATION FOR SEQ ID NO: 80:

5 (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1563 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

10

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 80:

	AATTCGGCAC	GAGNCAGAAA	CCTGCGGAAA	ATGGTAGCGA	TGGCGGCTGG	GCCGAGTGGG	60
15	TGTCTGGTGC	CGGCGTTTGG	GCTACGGTTG	TTGTTGGCGA	CTGTGCTTCA	AGCGGTGTCT	120
	GCTTTTGGGG	CAGAGTTTTC	ATCGGAGGCA	TGCAGAGAGT	TAGGCTTTTC	TAGCAACTTG	180
20	CTTTGCAGCT	CTTGTGATCT	TCTCGGACAG	TTCAACCTGC	TTCAGCTGGA	TCCTGATTGC	240
	AGAGGATGCT	GTCAGGAGGA	AGCACAATTT	GAAACCAAAA	AGCTGTATGC	AGGAGCTATT	300
	CTTGAAGTTT	GTGGATGAAA	ATTGGGAAGG	TTCCCTCAAG	TCCAAGCTTT	TGTTAGGAGT	360
25	GATAAACCCA	AACTGTTCAG	AGGACTGCAA	ATCAAGTATG	TCCGTGGTTC	AGACCCTGTA	420
	TTAAAGCTTT	TGGACGACAA	TGGGAACATT	GCTGAAGAAC	TGAGCATTCT	CAAATGGAAC	480
30	ACAGACAGTG	TAGAAGAATT	CCTGAGTGAA	AAGTTGGAAC	GCATATAAAT	CTTGCTTAAA	540
	TTTTGTCCTA	TCCTTTTGTT	ACCTTATCAA	ATGAAATATT	ACAGCACCTA	GAAAATAATT	600
	TAGTTTTGCT	TGCTTCCATT	GATCAGTCTT	TTACTTGAGG	CATTAAATAT	CTAATTAAAT	660
35	CGTGAAATGG	CAGTATAGTC	CATGATATCT	AAGGAGTTGG	CAAGCTTAAC	AAAACCCATT	720
	TTTTATAAAT	GTCCATCCTC	CTGCATTTGT	TGATACCACT	AACAAAATGC	TTTGTAACAG	780
40	ACTTGCGGTT	AATTATGCAA	ATGATAGTTT	GTGATAATTG	GTCCAGTTTT	ACGAACAACA	840
	GATTTCTAAA	TTAGAGAGGT	TAACAAGACA	GATGATTACT	ATGCCTCATG	TGCTGTGTGC	900
	TCTTTGAAAG	GAATGACAGC	AGACTACAAA	. GCAAATAAGA	TATACTGAGC	CTCAACAGAT	960
45	TGCCTGCTCC	TCAGAGTCTC	TCCTATTITT	GTATTACCCA	GCTTTCTTTT	TAATACAAAT	1020
	GTTATTTATA	GTTTACAATG	AATGCACTGC	ATAAAAACTT	TGTAGCTTCA	TTATTGTAAA	_1080
50	ACATATTCAA	GATCCTACAG	TAAGAGTGAA	ACATTCACAA	AGATTTGCGT	TAATGAAGAC	1140
	TACACAGAAA	ACCTTTCTAG	GGATTTGTGT	GGATCAGATA	CATACTTGGC	AAATTTTTGA	1200
	GTTTTACATT	CTTACAGAAA	AGTCCATTTA	AAAGTGATCA	A TTTGTAAGAC	CAAAATATAA	1260
55	ATAAAAAGTT	TCAAAAATCT	ATCTGAATTI	GGAATTCTTC	CIGGIIIGIIC	TTTCATGTTT	1320
	AAAAATGATG	TTTTTCAATG	CATTTTTTTC	ATGTAAGCCC	TTTTTTAGG	CAAAATGTAA	1380
60	AAATGGCTGT	' AATATTTAAA	ACTTATAACA	A TCTTATTGTT	r ggtaatagto	CTTTATATTT	1440

	GICIGATITI	ATTTTCAAA	GITTITICAT	TTATGAACAC	ATTTTCATTG	GTATATTATT	1500
	TAAGGAATAT	CTCTTGATAT	AGAATTTTTA	ТАТТАААААТ	GATTTTTCTT	TGCTTAAAAA	1560
5	AAA						1563

10 (2) INFORMATION FOR SEQ ID NO: 81:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1020 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 81:

TGCACGCTGG	CCATGTGGGN	GTTGGGCCAC	TGCGACCCCC	GGCGCTGCAC	GGGCCGCAAG	60
CTGGCCCGCC	TGGGGCTGGT	GCGCTGCCTG	CGCCTGGGCC	ACAGATTCGG	CGGTCTGGTG	120
CTGAGCCCCG	TGGGCAAGCA	GTACGCGTCC	CCCGCAGACA	GACAGCTGGT	GGCGCAGTCT	180
GGGGTCGCCG	TCATCGACTG	CTCCTGGGCC	AGGCTGGACG	AGACACCGTT	TGGGAAGATG	240
CGAGGGAGCC	ACTIGCGCCT	GTTGCCCTAC	CTCGTGGCCG	CCAACCCCGT	GAACTATGGC	300
CGGCCCTACA	GACTTTCCTG	CGTGGAAGCG	TTTGCTGCCA	CCTTCTGCAT	CGTAGGCTTT	360
CCAGACCTTG	CTGTCATTTT	GCTGCGGAAG	TTTAAATGGG	GCAAGGGCTT	CTTGGACCTG	420
AACCGCCAGC	TCCTGGACAA	GTACGCGGCC	TGCGGCAGCC	CGGAGGAGGT	GCTGCAGGCG	480
GAGCAGGAGT	TCTTGGCCAA	TGCCAAGGAG	AGCCCCCAGG	AGGAGGAGAT	CGATCCCTTC	540
GATGTGGATT	CAGGGAGAGA	GTTTGGAAAC	CCCAACAGGC	CTGTGGCCAG	CACCCGGCTG	600
CCCTCGGACA	CTGATGACAG	TGATGCGTCT	GAGGACCCAG	GGCCTKGCGC	CGAGCGCGGA	660
GGAGCCAGCA	GCAGCTGCTG	TGAAGAGGAG	CAGACGCAGG	GACGGGGGGC	TGAGGCCAGG	720
GCCCCGGCTG	AGGTTTGGAA	AGGAATCAAG	AAACGGCAGA	GAGACTGAGG	GTTGCAGACA	780
CATATATTTT	TGAGGCTGGG	TGACGAGAAA	ATCTAGAGAC	ATGAGGGACA	TAAATGGGCC	840
TGGCAGCCTC	GGCTCTTTGC	GGCTGCTGGC	AGGACTGAGC	TGTCCGGGTT	CTCCCCACAC	900
TTCCAGCACA	GCTGTGCTCT	GTGTCCTGCC	TCGGCGCTCT	CGCAAATGAA	GCTGCAGGCC	960
AAGAAAAAA	ААААААААА	АААААААА	аааааааа	АААААААА	GGGGGGGGC	1020
	CTGGCCCGCC CTGAGCCCCG GGGGTCGCCG GGGGTCGCCG CGAGGGAGCC CGGCCCTACA CCAGACCTTG AACCGCCAGC GAGCAGGAGT GATGTGGATT CCCTCGGACA GGAGCCAGCA GCCCCGGCTG CATATATTTT TGGCAGCCTC TTCCAGCACA	CTGGCCCGCC TGGGGCTGGT CTGAGCCCCG TGGGCAAGCA GGGGTCGCCG TCATCGACTG CGAGGGAGCC ACTTGCGCCT CGGCCCTACA GACTTTCCTG CCAGACCTTG CTGTCATTTT AACCGCCAGC TCCTGGACAA GAGCAGGAGT TCTTGGCCAA GATGTGGATT CAGGGAGAGA CCCTCGGACA CTGATGACAG GGAGCCAGCA GCAGCTGCTG GCCCCCGGCTG AGGTTTGGAA CATATATTTT TGAGGCTGGG TGGCAGCCTC GGCTCTTTGC TTCCAGCACA GCTGTGCTCT	CTGGCCCGCC TGGGCTGGT GCGCTGCCTG CTGAGCCCCG TGGCCAAGCA GTACGCGTCC GGGGTCGCCG TCATCGACTG CTCCTGGGCC CGAGGGAGCC ACTTGCGCCT GTTGCCCTAC CGGCCCTACA GACTTTCCTG CGTGGAAGCG CCAGACCTTG CTGTCATTTT GCTGCGGAAG AACCGCCAGC TCCTGGACAA GTACGCGGCC GAGCAGGAGT TCTTGGCCAA TGCCAAGGAG GATGTGGATT CAGGGAGAGA GTTTGGAAAC CCCTCGGACA CTGATGACAG TGATGCGTCT GGAGCCAGCA GCAGCTGCTG TGAAGAGGAG GCCCCGGCTG AGGTTTGGAA AGGAATCAAG CATATATTTT TGAGGCTGGG TGACGAGAAA TGGCAGCCTC GGCTCTTTGC GGCTGCTGCC	CTGGCCCGCC TGGGGCTGGT GCGCTGCCTG CGCCTGGGCC CTGAGCCCCG TGGGCAAGCA GTACGCGTCC CCCGCAGACA GGGGTCGCCG TCATCGACTG CTCCTGGGCC AGGCTGGACG CGAGGGAGCC ACTTGCGCCT GTTGCCCTAC CTGGTGGCCG CGGCCCTACA GACTTTCCTG CGTGGAAGCG TTTAAATGGG CCAGACCTTG CTGTCATTTT GCTGCGAAG TTTAAATGGG AACCGCCAGC TCCTGGACAA GTACGCGGCC TGCGGCAGCC GAGCAGGAGT TCTTGGCCAA TGCCAAGGAG AGCCCCCAGG GATGTGGATT CAGGGAGAG GTTTGGAAAC CCCAACAGGC CCCTCGGACA CTGATGACAG TGATGCGTCT GAGGACCCAG GGAGCCAGCA GCACCTGCTG TGAAGAGGAG CAGACGCAGG CCCCCGGCTG AGGTTTGGAA AGGAATCAAG AAACGGCAGA CATATATTTT TGAGGCTGGG TGACGAGAAA ATCTAGAGAC TGGCAGCCTC GGCTCTTTGC GGCTGCTGCC AGGACTGAGC TTCCAGCACA GCTGTGCTCT GTGTCCTGCC TCGGCGCTCT	CTGGCCCGCC TGGGCTGGT GCGCTGCCTG CGCCTGGGCC ACAGATTCGG CTGAGCCCCG TGGGCAAGCA GTACGCGTCC CCCGCAGACA GACAGCTGGT GGGGTCGCCG TCATCGACTG CTCCTGGGCC AGGCTGGACG AGACACCCGTT CGAGGGAGCC ACTTGCGCCT GTTGCCCTAC CTGGTGGCCG CCAACCCCGT CGGCCCTACA GACTTTCCTG CGTGGAAGCG TTTCCTGCCA CCTTCTGCAT CCAGACCTTG CTGTCATTTT GCTGCGGAAG TTTAAATGGG GCAAGGGCTT AACCGCCAGC TCCTGGACAA GTACGCGGCC TGCGGCAGCC CGGAGGAGGT GAGCAGGAGT TCTTGGCCAA TGCCAAGGAG AGCCCCCAGG AGGAGGAGT CCCTCGGACA CTGATGACAG TTTGGAAAC CCCAACAGGC CTGTGGCCAG CCCTCGGACA CTGATGACAG TGATGCGTCT GAGGACCCAG GGCCTKGCGC GGAGCCAGCA GCAGCTGCTG TGAAGAGGAG CAGACGCAGG GACGGGGGGC CCCCCGGCTG AGGTTTGGAA AGGAATCAAG AAACGGCAGA GAGACTGAGG CATATATTTT TGAGGCTGGG TGACGAGAAA ATCTAGAGAC ATGAGGGACA TGGCAGCCTC GGCTCTTTGC GGCTGCTGCC TCGGCGCTCT CGCCAAATGAA	TGCACGCTGG CCATGTGGGN GTTGGGCCAC TGCGACCCCC GGCGCTGCAC GGGCCGCAAG CTGGCCCGCC TGGGGCTGGT GCGCTGCCTG CGCCTGGGCC ACAGATTCGG CGGTCTGGTG CTGAGCCCCG TGGGCAAGCA GTACGCGTCC CCCGCAGACA GACAGCTGGT GGCGCAGTCT GGGGTCGCCG TCATCGACTG CTCCTGGGCC AGGCTGGACG AGACACCCGTT TGGGAAGATG CGAGGGAGCC ACTTGCGCCT GTTGCCCTAC CTGGTGGCCG CCAACCCCGT GAACTATGGC CGGCCCTACA GACTTTCCTG CGTGGAAGCG TTTGCTGCCA CCTTCTGCAT CGTAGGCTTT CCAGACCTTG CTGTCATTTT GCTGCGGAAG TTTAAATGGG GCAAGGGCTT CTTGGACCTG AACCGCCAGC TCCTGGACAA GTACGCGGCC TGCGGCAGCC CGGAGGAGGT GCTGCAGGCG GAGCAGGACT TCTTGGCCAA TGCCAAGGAG AGCCCCCAGG AGGAGGAGAT CGATCCCTTC GATGTGGATT CAGCGAGAGA GTTTGGAAAC CCCAACAGGC CTGTGGCCAG CACCCGGCTG CCCTCGGACA CTGATGACAG TGATGCGTCT GAGGACCAG GGCCTKGCGC CGAGCGCGGA GGAGCCAGCA GCACTCCTG TGAAGAGGAG CAGACCCAG GACGGGGGGC TGAGGCCAGG GCCCCCGGCTG AGGTTTGGAA AGGAATCAAG AAACGGCAGA GAGACTGAGG GTTGCAGGAC CATATATTTT TGAGGCTGGG TGACGAGAAA ATCTAGAGAC ATGAGGGACA TAAATGGGCC TGGCAGCCTC GGCTCTTTGC GGCTGCTGCC AGGACTGAGC TGTCCCGGGTT CTCCCCCACAC TTCCAGCACA GCTGTGCTCT GTGTCCTGCC TCGGCGCTCT CGCAAATGAA GCTGCAGGCC AAGAAAAAAA AAAAAAAAA AAAAAAAAA AAAAAAAA

55

(2) INFORMATION FOR SEQ ID NO: 82:

(i) SEQUENCE CHARACTERISTICS:

60 (A) LENGTH: 770 base pairs

480

232

	(B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 82:	
	TCGACCCACG CGTCCGGGCC GCCGTAGCGC GTCTTGGGTC TCCCGGCTGC CGCTGCTGCC	60
10	GCCGCCGCCT CGGGTCGTGG AGCCAGGAGC GACGTCACCG CCATGGCAGG CATCAAAGCT	120
	TIGATTAGTT TGTCCTTTGG AGGAGCAATC GGACTGATGT TTTTRATGCT TGGATGTGCC	180
	CTTCCAATAT ACAACAAATA CTGGCCCCTC TTTGTTCTAT TTTTTTACAT CCTTTCACCT	240
15	ATTCCATACT GCATAGCAAG AAGATTAGTG GATGATACAG ATGCTATGAG TAACGCTTGT	300
	AAGGAACTTG CCATCTTTCT TACAACGGGC ATTGTCGTGT CAGCTTTTGG ACTCCCTATT	360
20	GTATTTGCCA GAGCACATCT GATTGAGTGG GGAGCTTGTG CACTTGTTCT CACAGGAAAC	420
	ACAGTCATCT TTGCAACTAT ACTAGGCTTT TTCTTGGTCT TTGGAAGCAA TGACGACTTC	480
	AGCTGGCAGC AGTGGTGAAA AGAAATTACT GAACTATTGT CAAATGGACT TCCTGTCATT	540
25	TGTTGGCCAT TCACGCACAC AGGAGATGGG GCAGTTAATG CTGAATGGTA TAGCAAGCCT	600
	CTTGGGGGTA TTTTAGGTGC TCCCTTCTCA CTTTTATTGT AAGCATACTA TTTTCACAGA	660
30	GACTTGCTGA AGGATTAAAA GGATTTTCTC TTTTGGAAAA AAAAAAAAA AAAAACYCGA	720
	GGGGGGCCC GTWCCCATTC SCCCYATATG AATTCCNTTT TTACAATCCC	77
35	(2) INFORMATION FOR SEQ ID NO: 83:	
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 481 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 83:	
7.5	GAATTCGGCA CGAGCATAGT GTTAACCACT AGAATTCACT GCCCTTCCTA TCCAAAAATG	6
	ACACTACTGA TCATTTTTCT TCCTTTTSCT TTTACAACAT TMACAAATTC AGGTGGCTCT	12
50	TTCCCAGTAC GGTAGGCTGA TTCGTATGGA TGCACCACGG TTGGTGACTC CCCCCACCCC	18
	ACAGAGTTTC TGGCGTTCAT TCGGTTGAAC CCAAGGCCAG CAAGGGCTGA CTGGGAACAA	24
55	ACCGAACACT AGGCCGTGAA CCAATCGTCT CTCCGTGCCC GGGAGCGAMC CCGGGGGCCT	30
- -	TTCACTCTCC CAAGGACTCC ANGGGGGGGC CGGGTACCCA ATTCCGCCCC TATAGTGAAT	36
	CCCTNATTAC AATTCCACATT CCCCCGTCN TTTTTAACAAA GCTTTGGGTTTC AACTCCCAAA	40

AACCCCTTGG CGGTTTACCC CAACTTTAAT CCGCCTTTGC AAGCACATCC CCCCCTTTT

	c .	481
5		
	(2) INFORMATION FOR SEQ ID NO: 84:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 644 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 84:	
	GCTGGGATAG AGCATGAAAG GAGAACTGCT CCCTTTTCTG TTTCTCACAG TTTGGTTATG	60
20	GCTTTATAAA CTTKTATTTG GTGAAAGCCC CAGATACCCA AATGTCATTG GCAAAACTTA	120
20	TTTTTTTTTC TGGACAGATC AGATTTCTAG AGAGAGCAGA TTTCTAGAGA GATTAGCATT	180
	CATAGTAAGT GAAAATTGTC TAATTTTTTT AATCCATGCT ATTACTGGGC AGTAGGTCTA	240
25	ATTTTTTTTG ACAAAAATA GATCTATTTT CCTTATATAT TGATTTAGAA TCTTAAGTTA	300
	GAATTTTATA GAAGAAATGT CTGAGCAGTT CTATGTATGG AGGAGCAATT CAGCTTTTCA	360
30	GCAGCAACTT TATCTTTTGC CACTAGAGGG AGATCTGTGG TTGCTTTCTC CTTTGGAGAA	420
50	TAGCTGCTTT GCTTTTATTT TTAATTTCTA AGGTTGGAAT AGAACTTATT CTCAAAATTC	480
	CTTTAGTGTT ATTAAATATT TTCATTTATT AGTCAAAGGT AAGTTAATTA AGCTTGTTTA	540
35	ATGATGCCAA TCTTATGCTT TTCTGTAATC TTCAATTTTT AATAAATGTG AGTTAGATAC	600
	ТААСТСАААА ААААААААА ААААААААА ААААААААА	644
40		
	(2) INFORMATION FOR SEQ ID NO: 85:	
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1351 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 85:	
	GGCACGAGTG CGCASGCGTG GGGCTCTCTC CTTGTCAGTC GGCGCCGCGT GCGGGCTGGT	60
55	GGCTCTGTGG CAGCGGCGGC GGCAGGACTC CGGCACTATG AGCGGCTTCA GCACCGAGGA	120
JJ	GCGCGCCGCG CCNITCTCCC TGGAGTACCG AGTCTTCCTC AAAAATGAGA AAGGACAATA	180
	TATATCTCCA TTTCATGATA TTCCAATTTA TGCAGATAAG GATGTGTTTC ACATGGTAGT	240
60	TGAAGTACCA CGCTGGTCTA ATGCAAAAT GGAGATTCCT ACAAACGACC CTTTAAAACCA	200

300

	TATTAAACAA GATGTGAAAA AAGGAAAACT TCGCTATGTT GCGAATTTGT TCCCGTATAA	360
5	AGGATATATC TGGAACTATG GTGCCATCCC TCAGACTTGG GAAGACCCAG GGCACAATGA	420
,	TAAACATACT GGCTGTTGTG GTGACAATGA CCCAATTGAT GTGTGTGAAA TTGGAAGCAA	480
	GGTATGTGCA AGAGGTGAAA TAATTGGCGT GAAAGTTCTA GGCATATTGG CTATGATTGA	540
10	CGAAGGGGAA ACCGACTGGA AAGTCATTGC CATTAATGTG GATGATCCTG ATGCAGCCAA	600
	TTATAATGAT ATCAATGATG TCAAACGGCT GAAACCTGGC TACTTAGAAG CTACTGTGGA	660
15	CTGGTTTAGA AGGTATAAGG TTCCTGATGG AAAACCAGAA AATGAGTTTG CGTTTAATGC	720
13	AGAATITAAA GATAAGGACT TTGCCATTGA TATTATTAAA AGCACTCATG ACCATTGGAA	780
	AGCATTAGTG ACTAAGAAAA CGAATGGAAA AGGAATCAGT TGCATGAATA CAACTTTGTC	840
20	TGAGAGCCCC TTCAAGTGTG ATCCTGATGC TGCCAGAGCC ATTGTGGATG CTTTACCACC	900
	ACCCTGTGAA TCTGCCTGCA CAGTACCAAC AGACGTGGAT AAGTGGTTCC ATCACCAGAA	960
25	AAACTAATGA GATTTCTCTG GAATACAAGC TGATATTGCT ACATCGTGTT CATCTGGATG	1020
23	TATTAGAAGT AAAAGTAGTA GCTTTTCAAA GCTTTAAATT TGTAGAACTC ATCTAACTAA	1080
	AGTAAATTCT GCTGTGACTA ATCCAATATA CTCAGAATGT TATCCATCTA AAGCATTTTT	1140
30	CATATCTCAA CTAAGATAAC TTTTAGCACA TGCTTAAATA TCAAAGCAGT TGTCATTTGG	1200
	AAGTCACTTG TGAATAGATG TGCAAGGGGA GCACATATTG GATGTATATG TTACCATATG	1260
35	TTAGGAAATA AAATTATTTT GCTGAAAAAA AAAAAAAAA AACCNCGGGG GGGGCCCCGG	1320
55	TCCCCATTIG GCCCTTTGGG GGGNGGTTTT A	1351
40	(2) INFORMATION FOR SEQ ID NO: 86:	
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2527 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 86:	
50	CTCTTGCTAC CTTCCCGGCG CAGAGAACCC CGGCTGCTCA GCGCGCTCCG GGGTCATGGA	60
	GATCCCCGGG AGCCTGTGCA AGAAAGTCAA GCTGAGCAAT AACGCGCAGA ACTGGGGAAT	120
55	GCAGAGASCA ACCAATGTCA CCTACCAAGC CCATCATGTC AGCAGGAACA AGAGAGGTCA	180
	GOTTOCTIVESCE ACCEGEGGGGG COMMUNICAGE MYSCACACACTUR MESCURADOR COMMUNICAGE	240

AGCGGGAAAG ACTACTGTGA GCATGGCCTT GGAGGAGTAC CTGGTTTGTC ATGGTATTCC

	ATGCTACACT	CTGGATGGTG	ACAATATTCG	TCAAGGTCTC	AATAAAAATC	TTGGCTTTAG	360
	TCCTGAAGAC	AGAGAAGAGA	ATGTTCGACG	CATCGCAGAA	GTTGCTAAAC	TGTTTGCAGA	420
5	TGCTGGCTTA	GTGTGCATCA	CAAGTTTCAT	ATCACCTTAC	ACTCAGGATC	GCAACAATGC	480
	AAGGCAAATT	CATGAAGGTG	CAAGTTTACC	GTTTTTTGAA	GTATTTGTTĞ	ATGCTCCTCT	540
10	GCATGTTTGT	GAACAGAGGG	ATGTCAAAGG	ACTCTACAAA	AAAGCCCGGG	CAGGAGAAAT	600
10	TAAAGGTTTC	ACTGGGATCG	ATTCTGAATA	TGAAAAGCCA	GAGGCCCCTG	AGTTGGTGCT	660
	GAAAACAGAC	TCCTGTGATG	TAAATGACTG	TGTCCAGCAA	GTTGTGGAAC	TTCTACAGGA	720
15	ACGGGATATT	GTACCTGTGG	ATGCATCTTA	TGAAGTAAAA	GAACTATATG	TGCCAGAAAA	780
	TAAACTTCAT	TTGGCAAAAA	CAGATGCGGA	AACATTACCA	GCACTGAAAA	TTAATAAAGT	840
20	GGATATGCAG	TGGGTGCAGG	TTTTGGCAGA	AGGTTGGGCA	ACCCCATTGA	ATGGCTTTAT	900
	GAGAGAGAGG	GAGTACTTGC	AGTGCCTTCA	TTTTGATTGT	CTTCTGGATG	GAGGTGTCAT	960
	TAACTTGTCA	GTACCTATAG	TTCTGACTGC	GACTCATGAA	GATAAAGAGA	GGCTGGACGG	1020
25	CTGTACAGCA	TTTGCTCTGA	TGTATGAGGG	CCGCCGTGTG	GCCATTCTTC	GCAATCCAGA	1080
	GTTTTTTGAG	CACAGGAAAG	AGGAGCGCTG	TGCCAGACAG	TGGGGAACGA	CATGCAAGAA	1140
30	CCACCCCTAT	ATTAAGATGG	TGATGGAACA	AGGAGATTGG	CTGATTGGAG	GAGATCTTCA	1200
	AGTCTTGGAT	CGAGTTTATT	GGAATGATGG	TCTTGATCAG	TATCGTCTTA	CTCCTACTGA	1260
	GCTAAAGCAG	AAATTTAAAG	ATATGAATGC	TGATGCTGTC	TTTGCATTTC	AACTACGCAA	1320
35	CCCAGTGCAC	AATGGACATG	CCCTGTTAAT	GCAGGATACC	CATAAGCAAC	TTCTAGAGAG	1380
	GGGCTACCGG	CGCCCTGTCC	TCCTCCTCCA	CCCTCTGGGT	GGCTGGACAA	AGGATGACGA	1440
40	TGTTCCTTTG	ATGTGGCGTA	TGAAGCAGCA	TGCTGCAGTG	TTGGAGGAAG	GAGTTCTGAA	1500
. •	TCCTGAGACG	ACAGTGGTGG	CCATCTTCCC	ATCTCCCATG	ATGTATGCTG	GACCAACTGA	1560
	GGTCCAGTGG	CATTGCAGAG	CACGGATGGT	TGCAGGAGCC	AACTTTTACA	TTGTTGGACG	1620
45	AGACCCTGCT	GGCATGCCTC	ATCCAGAAAC	AGGGAAGGAT	CTTTATGAGC	CAAGTCATGG	1680
	TGCCAAAGTG	CTGACGATGG	CCCCTGGTTT	AATCACTTTG	GAAATAGTTC	CCTTTCGAGT	1740
50	TGCAGCTTAC	AACAAGAAAA	AGAAGCGTAT	GGACTACTAT	GACTCTGAAC	ACCATGAAGA	1800
	CTTTGAATTT	ATTTCAGGAA	CACGAATGCG	CAAACTTGCT	CGAGAAGGCC	AGAAACCACC	1860
	TGAAGGTTTC	ATGGCTCCCA	AGGCTTGGAC	CGTGCTGACA	GAATACTACA	AATCCTTGGA	1920
55	GAAAGCTTAG	GCTGTTAACC	CAGTCACTCC	ACCTTTGACA	CATTACTAGT	' AACAAGAGGG	1980
	GACCACATAG	TCTCTGTTGG	CATTTCTTTC	TGGTGTCTGT	CTGGACATGC	TTCCTAAAAA	2040
60	CAGACCATTT	TCCTTAACTT	GCATCAGTTI	TGGTCTGCCT	TATGAGTTCT	GTTTTGAACA	2100

960

	AGIGTAACAC ACTGATGGTT TTAATGTATC TTTTCCACTT ATTATAGTTA TATTCCTACA	2160
	ATACAATTTT AAAATTGTCT TTTTATATTA TATTTATGCT TCTGTGTCAT GATTTTTTCA	2220
5	AGCTGTTATA TTAGTTGTAA CCAGTAGTAT TCACATTAAA TCTTGCTTTT TTTCCCCTTA	2280
	AAAAAAGAAA AAAATTACCA AACAATAAAC TTGGCTAGAC CTTGTTTTGA GGATTTTACA	2340
10	AGACCTTTGT AGCGATTAGA TTTTTTTTCT ACATTGAAAA TAGAAACTGC TTCCTTTCTT	2400
10	CTTTCCAGTC AGCTATTGGT CTTTCCAGCT GTTATAATCT AAAGTATTCT TATGATCTGT	2460
	GTAAGCTCTG AATGAACTTC TTTACTCAAT AAAATTAATT TTTTGGCTTC TTAAAAAAAA	2520
15	AAAAAA	2527
20	(2) INFORMATION FOR SEO ID NO: 87:	
	(i) SEQUENCE CHARACTERISTICS:	
25	(A) LENGTH: 2566 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 87:	
30	CCCAAGAATT CGGCACGAGC GNGGCAWAAK TGGGATTTCT GAAACCTGTA GGCCCCAAGC	60
	CCATCAACTT GCCCAAAGAA GATTCCAAAC CTACATTTCC CTGGCCTSCT GGAAACAAGC	120
35	CATCTCTTCA CAGTGTAAAC CAAGACCATG ACTTAAAGCC ACTAGGCCGA AATCTGGGCC	180
	TACTCCTCCA ACCTCAGAAA ATGAACAGAA GCAAGCKTTT CCCAAATTGA CTGGGGTTAA	240
	AGGGAAATTT ATGTCAGCAT CACAAGATCT TGAACCCAAG CCCCTCTTCC CCAAACCCGC	300
40	CTTTGGCCAG AAGCCGCCCC TAAGTACCGA GAACTCCCAT GAAGACGAAA GCCCCATGAA	360
	GAATGTGTCT TCATCAAAAG GGTCCCCAGC TCCCCTGGGA GTCAGGTCCA AAAGCGGCCC	420
45	TTTAAAACCA GCAAGGGAAG ACTCAGAAAA TAAAGACCAT GCAGGGGAGA TTTCAAGTTT	480
	GCCCTTTCCT GGAGTGGTTT TGAAACCTGC TGCGAGCAGG GGAGGCCCAG GTCTCTCCAA	540
	AAATGGTGAA GAAAAAAAGG AAGATAGGAA GATAGATGCT GCTAAGAACA CCTTCCAGAG	600
50	CAAAATAAAT CAGGAAGAGT TGGCCTCAGG GACTCCTCCT GCCAGGTTCC CTAAGGCCCC	660
	TTCTAAGCTG ACAGTGGGG GGCCATGGGG CCAAAGTCAG GAAAAGGAAA AGGGAGACAA	720
55	GAATTCAGCC ACCCCGAAAC AGAAGCCATT GCCTCCCTTG TTTACCTTGG GTCCACCTCC	780
	ACCAAAACCC AACAGACCAC CAAATGTTGA CCTGACGAAA TTCCACAAAA CCTCTTCTGG	840
	AAACAGTACT AGCAAAGGCC AGACGTCTTA CTCAACAACT TCCCTGCCAC CACCTCCACC	900

ATCCCATCCG GCCAGCCAAC CACCATTGCC AGCATCTCAC CCATCACAAC CACCAGTCCC

	AAGCCTACCT	CCCAGAAACA	TTAAACCTCC	GTTTGACCTA	AAAAGCCCTG	TCAATGAAGA	1020
5	CAATCAAGAT	GGTGTCACGC	ACTCTGATGG	TGCTGGAAAT	CTAGATGAGG	AACAAGACAG	1080
	TGAAGGAGAA	ACATATGAAG	ACATAGAAGC	ATCCAAAGAA	AGAGAGAAGA	AAAGGGAAAA	1140
	GGAAGAAAAG	AAGAGGTTAG	AGCTGGAGAA	AAAGGAACAG	AAAGAGAAAG	AAAAGAAAGA	1200
10	ACAAGAAATA	AAGAAGAAAT	TTAAACTAAC	AGGCCCTATT	CAAGTCATCC	ATCTTGCAAA	1260
	AGCTTGTTGT	GATGTCAAAG	GAGGAAAGAA	TGAACTGAGC	TTCAAGCAAG	GAGAGCAAAT	1320
15	TGAAATCATC	CGCATCACAG	ACAACCCAGA	AGGAAAATGG	TTGGGCAGAA	CAGCAAGGGG	1380
	TTCATATGGC	ТАТАТТАААА	CAACTGCTGT	AGAGATTGAC	TATGATTCTT	TGAAACTGAA	1440
	AAAAGACTCT	CTTGGTGCCC	CTTCAAGACC	TATTGAAGAT	GACCAAGAAG	TATATGATGA	1500
20	TGTTGCAGAG	CAGGATGATA	TTAGCAGCCA	CAGTCAGAGT	GGAAGTGGAG	GGATATTCCC	1560
	TCCACCACCA	GATGATGACA	TTTATGATGG	GATTGAAGAG	GAAGATGCTG	ATGATGGCTC	1620
25	CACACTACAG	GTTCAAGAGA	AGAGTAATAC	GTGGTCCTGG	GGGATTTTGA	AGATGTTAAA	1680
	GGGAAAAGAT	GACAGAAAGA	AAAGTATACG	AGAGAAACCT	AAAGTCTCTG	ACTCAGACAA	1740
	TAATGAAGGT	TCATCTTTCC	CTGCTCCTCC	TAAACAATTG	GACATGGGAG	ATGAAGTTTA	1800
30	CGATGATGTG	GATACCTCTG	ATTICCCTGT	TTCATCAGCA	GAGATGAGTC	AAGGAACTAA	1860
	TGTTGGAAAA	GCTAAGACAG	AAGAAAAGGA	CCTTAAGAAG	CTAAAAAAGC	AGRAAAARA	1920
35	ARAAAAAGAC	TTCAGGAAAA	AATTTAAATA	TGATGGTGAA	ATTAGAGTCC	TATATTCAAC	1980
	TAAAGTTACA	ACTTCCATAA	СТТСТААААА	GTGGGGAACC	AGAGATCTAC	AGGTAAAACC	2040
	TGGTGAATCT	CTAGAAGTTA	TACAAACCAC	AGATGACACA	AAAGTTCTCT	GCAGAAATGA	2100
40	AGAAGGGAAA	TATGGTTATG	TCCTTCGGAG	TTACCTAGCG	GACAATGATG	GAGAGATCTA	2160
	TGATGATATT	GCTGATGGCT	GCATCTATGA	CAATGACTAG	CACTCAACTT	TGGTCATTCT	2220
45	GCTGTGTTCA	TTAGGTGCCA	ATGTGAAGTC	TGGATTTTAA	TTGGCATGTT	ATTGGGTATC	2280
	AAGAAAATTA	ATGCACAAAA	CCACTTATTA	TCATTTGTTA	TGAAATCCCA	ATTATCTTTA	2340
	CAAAGTGTTT	AAAGTTTGAA	CATAGAAAAT	AATCTCTCTG	CTTAATTGTT	ATCTCAGAAG	2400
50	ACTACATTAG	TGAGATGTAA	GAATTATTAA	ATATTCCATT	TCCGCTTTGG	CTACAATTAT	2460
	GAAGAAGTTG	AAGGTACTTC	TTTTAGACCA	CCAGTAAATA	ATCCTCCTTC	AAAAAAAA	2520
55	ÄATAAAAAAA	ААААААААА	ACTCGAGGGG	GGGCCCGGTA	CCCAAT		2566

⁽²⁾ INFORMATION FOR SEQ ID NO: 88:

60

्रेस प्रतिकार के एक के मिना प्रतिकार के और कुछान प्रतिकार के लिए का पान के लिए में को के की कार के का का का का कार्य

5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 540 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 88:	
10	GAATTCGGCA CGAGGCTTTC TGTGTCCTCT GTGGCTGCTT TAGTGTGCCA CCAGGGGCAG	60
••	ACTTGGGTGG GTTGCAGCAG AGATGGCATG GCCCTCAAGG TCCAAGATGT TTACTCTCTT	120
	GCCGGTCCTC TGTTATCTCT GGTCTTTGTG GTTGCCACAG TTTTCTTGGA TCCAGGAGTT	180
15	AAAGGCAGTC CTGAGGGATG ATGGCCTCAT CTCCGCAGTT GCYTGGAATG CTGAATTTCA	240
	GACGTGCTAA AGGAGGGTTG CAGACATTGT GTGGWATGCA TTCAGACCCC AGATGTGGGT	300
20	GCAGGAAGGC AGGCATGGCA CAGCCAGGTA GAGACTGGTT TCCAGGCCCA AGCAGCCTȚC	360
20	AGCAGCTGTG CGCCTTGTTT CTGATGTTGT TTGGGAGTAA GAATAATGTA GACATGGGGG	420
	GTCATGARGC TCAATAAAAA CTTCAAGGAA ACCTCCCATG GCATGGTTGG GCGCAGTGAC	480
25	TCATGCCTGT AACCCCAGCA CTGTGGAATG CCAAGGTGGA AGGATCGCTT GAGGCCAAGA	540
		•
30	(2) INFORMATION FOR SEQ ID NO: 89:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1863 base pairs	
35	(B) TYPE: nucleic àcid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89:	
40	TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT	60
	CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG	120
45	CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCG CCCTTCGAGG GCGCCCCAGG	180
40	CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA	240
	CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG	300
50	CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG	360
	CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA	420

TTTTGCACTT CAACCAGATG ACGTGTACTA CTGTGGAATA AAGTACATCA AAGATGATGT

CATCTTAAAT GAGCCCTCTG CAGATGCCCC AGCTGCTCTC TACCAGACAA TTGAAGAAAA

TATTAAAATC TTTGAAGAAG AAGAAGTTGA ATTTATCAGT GTGCCTGTCC CAGAGTTTGC

AGATAGTGAT CCTGCCAACA TTGTTCATGA CTTTAACAAG AAACTTACAG CCTATTTAGA



480

540

600

TCTTAACCTG GATAAGTGCT ATGTGATCCC TCTGAACACT TCCATTGTTA TGCCACCCAG

720

5	AAACCTACTG GAGTTACTTA TTAACATCAA GGCTGGAACC TATTTGCCTC AGTCCTATCT	780
,	GATTCATGAG CACATGGTTA TTACTGATCG CATTGAAAAC ATTGATCACC TGGGTTTCTT	840
	TATTTATCGA CTGTGTCATG ACAAGGAAAC TTACAAACTG CAACGCAGAG AAACTATTAA	900
10	AGGTATTCAG AAACGTGAAG CCAGCAATTG TTTCGCAATT CGGCATTTTG AAAACAAATT	960
	TGCCGTGGAA ACTTTAATTT GTTCTTGAAC AGTCAAGAAA AACATTATTG AGGAAAATTA	1020
15	ATATCACAGC ATAACCCCAC CCTTTACATT TTGTGCAGTG ATTATTTTTT AAAGTCTTCT	1080
13	TTCATGTAAG TAGCAAACAG GGCTTTACTA TCTTTTCATC TCATTAATTC AATTAAAACC	1140
	ATTACCTTAA AATTTTTTTC TTTCGAAGTG TGGTGTCTTT TATATTTGAA TTAGTAACTG	1200
20	TATGAAGTCA TAGATAATAG TACATGTCAC CTTAGGTAGT AGGAAGAATT ACAATTTCTT	1260
	TAAATCATTT ATCTGGATTT TTATGTTTTA TTAGCATTTT CAAGAAGACG GATTATCTAG	1320
25	AGAATAATCA TATATATGCA TACGTAAAAA TGGACCACAG TGACTTATTT GTAGTTGTTA	1380
23	GTTGCCCTGC TACCTAGTTT GTTAGTGCAT TTGAGCACAC ATTTTAATTT TCCTCTAATT	1440
	AAAATGTGCA GTATTTTCAG TGTCAAATAT ATTTAACTAT TTAGAGAATG ATTTCCACCT	1500
30	TTATGTTTTA ATATCCTAGG CATCTGCTGT AATAATATTT TAGAAAATGT TTGGAATTTA	1560
	AGAAATAACT TGTGTTACTA ATTTGTATAA CCCATATCTG TGCAATGGAA TATAAATATC	1620
35	ACAAAGTTGT TTAACTAGAC TGCGTGTTGT TTTTCCCGTA TAATAAAACC AAAGAATAGT	1680
	TTGGTTCTTC AAATCTTAAG AGAATCCACA TAAAAGAAGA AACTATTTTT TAAAAATTCA	1740
	CTTCTATATA TACAATGAGT AAAATCACAG ATTTTTCTT TAAATAAAAA TAAGTCATTT	1800
40	TAATAACTAA ACCAGATTCT TTGTGATACT ATTAANGTAA CATTTAGCCC CAAAAAAAAA	1860
	AAA	1863
45		
	(2) INFORMATION FOR SEQ ID NO: 90:	
	(i) SEOUENCE CHARACTERISTICS:	
50	(A) LENGTH: 2478 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 90:	
	GGCACAGCGG CACGAGGTGA GCTGAGCCGG TGGGTGAGCG GCGGCCACGG CATCCTGTGC	60
	THE COLOREST ACCACCAAAC ATTENDED CATTER CATTER COACACTOTTE TO THE TOTAL COLOREST COACACTOTTE TO THE TOTAL COLOREST COLOR	120

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	GTCCCTGTGC ACAGCCTTTG CCTTGAGCAA ACCCACAGAA AAGAAGGACC GTGTACATCA	180
	TGAGCCTCAG CTCAGTGACA AGGTTCACAA TGATGCTCAG AGTTTTGATT ATGACCATGA	240
5	TGCCTTCTTG GGTGCTGAAG AAGCAAAGAC CTTTGATCAG CTGACACCAG AAGAGAGCAA	300
	GGAAAGGCTT GGAAAGATTG TAAGTAAAAT AGATGGCGAC AAGGACGGGT TTGTCACTGT	360
10	GGATGAGCTC AAAGACTGGA TTAAATTTGC ACAAAAGCGC TGGATTTACG AGGATGTAGA	420
10	GCGACAGTGG AAGGGGCATG ACCTCAATGA GGACGGCCTC GTTTCCTGGG AGGAGTATAA	480
	AAATGCCACC TACGGCTACG TTTTAGATGA TCCAGATCCT GATGATGGAT TTAACTATAA	.540
15	ACAGATGATG GTTAGAGATG AGCGGAGGTT TAAAATGGCA GACAAGGATG GAGACCTCAT	600
	TGCCACCAAG GAGGAGTTCA CAGCTTTCCT GCACCCTGAG GAGTATGACT ACATGAAAGA	660
20	TATAGTAGTA CAGGAAACAA TGGAAGATAT AGATAAGAAT GCTGATGGTT TCATTGATCT	720
20	AGAAGAGTAT ATTGGTGACA TGTACAGCCA TGATGGGAAT ACTGATGAGC CAGAATGGGT	780
	AAAGACAGAG CGAGAGCAGT TTGTTGAGTT TCGGGATAAG AACCGTGATG GGAAGATGGA	840
25	CAAGGAAGAG ACCAAAGACT GGATCCTTCC CTCAGACTAT GATCATGCAG AGGCAGAAGC	900
	CAGGCACCTG GTCTATGAAT CAGACCAAAA CAAGGATGGC AAGCTTACCA AGGAGGAGAT	960
30	CGTTGACAAG TATGACTTAT TTGTTGGCAG CCAGGCCACA GATTTTGGGG AGGCCTTAGT	1020
30	ACGGCATGAT GAGTTCTGAG CTRCGGAGGA ACCCTCATTT CCTCAAAAGT AATTTATTTT	1080
	TACAGCTTCT GGTTTCACAT GAAATTGTTT GCGCTACTGA GACTGTTACT ACAAACTTTT	1140
35	TAAGACATGA AAAGGCGTAA TGAAAACCAT CCCGTCCCCA TTCCTCCTCC TCTCTGAGGG	1200
,	ACTGGAGGGA AGCCGTGCTT CTGAGGAACA ACTCTAATTA GTACACTTGT GTTTGTAGAT	1260
40	TTACACTTTG TATTATGTAT TAACATGGCG TGTTTATTTT TGTATTTTTC TCTGGTTGGG	1320
40	AGTATGATAT GAAGGATCAA GATCCTCAAC TCACACATGT AGACAAACAT TAGCTCTTTA	1380
	CTCTTTCTCA ACCCCTTTTA TGATTTTAAT AATTCTCACT TAACTAATTT TGTAAGCCTG	1440
45	AGATCAATAA GAAATGTTCA GGAGAGAGGA AAGAAAAAAA ATATATGCTC CACAATTTAT	1500
	ATTITAGAGAG AGAACACTITA GICTIGCCIG TCAAAAAGIC CAACATITCA TAGGIAGIAG	1560
	GGGCCACATA TTACATTCAG TTGCTATAGG TCCAGCAACT GAACCTGCCA TTACCTGGGC	1620
50	AAGGAAAGAT CCCTTTGCTC TAGGAAAGCT TGGCCCAAAT TGATTTTCTT CTTTTTCCCC	168
	CTGTAGGACT GACTGTTGGC TAATTTTGTC AAGCACAGCT GTGGTGGGAA GAGTTAGGGC	174
55	CAGTGTCTTG AAAATCAATC AAGTAGTGAA TGTGATCTCT TTGCAGAGCT ATAGATAGAA	180
	ACAGCTGGAA AACTAAAGGA AAAATACAAG TGTTTTCGGG GCATACATTT TTTTTCTGGG	186
60	TGTGCATCTG TTGAAATGCT CAAGACTTAA TTATTTGCCT TTTGAAATCA CTGTAAATGC	192

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	CCCCATCCGG	TTCCTCTTCT	TCCCAGGTGT	GCCAAGGAAT	TAATCTTGGT	TTCACTACAA	1980
	TTAAAATTCA	CTCCTTTCCA	ATCATGTCAT	TGAAAGTGCC	TTTAACGAAA	GAAATGGTCA	2040
5	CTGAATGGGA	ATTCTCTTAA	GAAACCCTGA	GATTAAAAAA	AGACTATTTG	GATAACTTAT	2100
	AGGAAAGCCT	AGAACCTCCC	AGTAGAGTGG	GGATTTTTTT	CTTCTTCCCT	TTCTCTTTTG	2160
10	GACAATAGTT	AAATTAGCAG	TATTAGTTAT	GAGTTTGGTT	GCAGTGTTCT	TATCTTGTGG	2220
	GCTGATTTCC	AAAAACCACA	TGCTGCTGAA	TTTACCAGGG	ATCCTCATAC	CTCACAATGC	2280
	AAACCACTTA	CTACCAGGCC	TTTTTCTGTG	TCCACTGGAG	AGCTTGAGCT	CACACTCAAA	2340
15	GATCAGAGGA	CCTACAGAGA	GGGCTCTTTG	GTTTGAGGAC	CATGGCTTAC	CTTTCCTGCC	2400
	TTTGACCCAT	CACACCCCAT	TTCCTCCTCT	TTCCCTCTCC	CCGCTGCCAA	TTCCTGCAGC	2460
20	CCGGGGGAAC	CACTAGTT					2478

(2) INFORMATION FOR SEQ ID NO: 91:

25

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2058 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- 30 (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 91:

35	TCGGCCTTGC	TTTTGTGGYC	TTCCTCTGTG	GCCAGAGCGT	TTTCATCACC	AAGCCTCCTG	60
	ATGGCAGTNC	CTTCACCGAT	ATGTTCAAGA	TACTGACGTA	TTCCTGCTGT	TCCCAGAAGC	120
	GAAGTGGAGA	GCGCCAGAGT	AATGGTGAAG	GCATTGGAGT	NTTTCAGCAA	TCTTCTAAAC	180
40	AAAGTCTGTT	TGATTCATGT	AAGATGTCTC	ATGGTGGGCC	ATTTACAGAA	GAGAAAGTGG	240
	AAGATGTGAA	AGCTCTGGTC	AAGATTGTCC	CTGTTTTCTT	GGCTTTGATA	CCTTACTGGA	300
45	CAGTGTATTT	CCAAATGCAG	ACAACATATG	TTTTACAGAG	TCTTCATTTG	AGGATTCCAG	360
	AAATTTCAAA	TATTACAACC	ACTCCTCACA	CGCTCCCTGC	AGCCTGGCTG	ACCATGTTTG	420
	ATGCTGTGCT	CATCCTCCTG	CTCATCCCTC	TGAAGGACAA	ACTGGTCGAT	CCCATTTTGA	480
50	GAAGACATGG	CCTGCTCCCA	TCCTCCCTGA	AGAGGATCGC	CGTGGGCATG	TTCTTTGTCA	540
	TGTGCTCRGC	CTTTGCTGCA	GGAATTTTGG	AGAGTAAAAG	GCTGAACCTT	GTTAAAGAGA	600
55	AAACCÄTTAA	TCAGACCATC	GGCAACGTCG	TCTACCATGC	TGCCGATCTG	TCGCTGTGGT	660
	GGCAGGTGCC	GCAGTACTTG	CTGATTGGGA	TCAGCGAGAT	CTTTGCAAGT	ATCGCAGGCC	720
	TGGAATTTGC	ATACTCAGCT	GCCCCCAAGT	CCATGCAGAG	TGCCATAATG	GGCTTGTTCT	780
60	TTTTCTCTC	TGGCGTCGGG	TCGTTCGTGG	GTTCTGGACT	GCTGGCACTG	GTGTCTATCA	840

	AAGCCATCGG	ATGGATGAGC	AGTCACACAG	ACTTTGGTAA	TATTAACGGC	TGCTATTTGA	900
_	ACTATTACTT	TTTCCTTCTG	GCTGCTATTC	AAGGAGCTAC	CCTCCTGCTT	TTCCTCATTA	960
5	TTTCTGTGAA	ATATGACCAT	CATCGAGACC	ATCAGCGATC	AAGAGCCAAT	GGCGTGCCCA	1020
	CCAGCAGGAG	GGCCTGACCT	TCCTGAGGCC	ATGTGCGGTT	TCTGAGGCTG	ACATGTCAGT	1080
10	AACTGACTGG	GGTGCACTGA	GAACAGGCAA	GACTTTAAAT	TCCCATAAAA	TGTCTGACTT	1140
	CACTGAAACT	TGCATGTTGC	CTGGATTGAT	TTCTTCTTTC	CCTCTATCCA	AAGGAGCTTG	1200
15	GTAAGTGCCT	TACTGCAGCG	TGTCTCCTGG	CACGCTGGGC	CCTCCGGGAG	GAGAGCTGCA	1260
13	GATTTCGAGT	ATGTCGCTTG	TCATTCAAGG	TCTCTGTGAA	TCCTCTAGCT	GGGTTCCCTT	1320
	TTTTACAGAA	ACTCACAAAT	GGAGATTGCA	AAGTCTTGGG	GAACTCCACG	TGTTAGTTGG	1380
20	CATCCCAGTT	TCTTAAACAA	ATAGTATCAC	CTGCTTCCCA	TAGCCATATC	TCACTGTAAA	1440
	AAAAAAATT	AATAAACTGT	TACTTATATT	TAAGAAAGTG	AGGATTTTT	TTTTTTAAAG	1500
25	ATAAAAGCAT	GGTCAGATGC	TGCAAGGATT	ттасатааат	GCCATATTTA	TGGTTTCCTT	1560
<u> </u>	CCTGAGAACA	ATCTTGCTCT	TGCCATGTTC	TTTGATTTAG	GCTGGTAGTA	AACACATTTC	1620
	ATCTGCTGCT	TCAAAAAGTA	CTTACTTTT	AAACCATCAA	CATTACTTTT	CTTTCTTAAG	1680
30	GCAAGGCATG	CATAAGAGTC	ATTTGAGACC	ATGTGTCCCA	TCTCAAGCCA	CAGAGCAACT	1740
	CACGGGGTAC	TTCACACCTT	ACCTAGTCAG	AGTGCTTATA	TATAGCTTTA	TTTTGGTACG	1800
35	ATTGAGACTA	AAGACTGATC	ATGGTTGTAT	GTAAGGAAAA	CATTCTTTTG	AACAGAAATA	1860
<i>J J</i>	GTGTAATTAA	AAATAATTGA	. AAGTGTTAAA	TGTGAACTTG	AGCTGTTTGA	CCAGTCACAT	1920
	TTTTGTATTG	TTACTGTACG	TGTATCTGGG	GCTTCTCCGT	' TTGTTAATAC	TTTTTCTGTA	1980
40	TTTGTTGCTG	TATTITIGGO	ATAACTTTAT	TATAAAAAGC	ATCTCAAATC	CGAAAWAAAA	2040
	AAAAAAAA	AAAAAAAC					2058
45							
	(2) INFORM	MATION FOR S	SEO ID NO: 9	92:			

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1411 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

55 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 92:

> GGCACAGGAG CGACCCGGGA GAAGGAGGGC CAMGAKGCGG AAGCCGGAGGA GTCTCCAGGA 60 GACCCGGGGA CAGCATCGCC CAGGCCCCTG TTTGCAGGCC TTTCAGATAT ATCCATCTCA

	CAAGACATCC	CCGTAGAAGG	AGAAATCACC	ATTCCTATGA	GATCTCGCAT	CCGGGAGTTT	180
	GACAGCTCCA	CATTAAATGA	ATCTGTTCGC	AATACCATCA	TGCGTGATCT	AAAAGCTGTT	240
5	GGGAAAAAT	TCATGCATGT	TTTGTACCCA	AGGAAAAGTA	ATACTCTTTT	GAGAGATTGG	300
	GATTTGTGGG	GCCCTTTGAT	CCTTTGTGTG	ACACTCGCAT	TAATGCTGCA	AAGAGACTCT	360
10	GCAGATAGTG	AAAAAGATGG	AGGGCCCCAA	TTTGCAGAGG	TGTTTGTCAT	TGTCTGGTTT	420
10	GGTGCAGTTA	CCATCACCCT	CAACTCAAAA	CTTCTTGGAG	GGAACATATC	TTTTTTCAG	480
	AGCCTCTGTG	TGCTGGGTTA	CTGTATACTT	CCCTTGACAG	TAGCAATGCT	GATTTGCCGG	540
15	CTGGTACTTT	TGGCTGATCC	AGGACCTGTA	AACTTCATGG	TTCGGCTTTT	TGTGGTGATT	600
	GTGATGTTTG	CCTGGTCTAT	AGTTGCCTCC	ACAGCTTTCC	TTGCTGATAG	CCAGCCTCCA	660
20	AACCGCAGAG	CCCTAGCTGT	TTATCCTGTT	TTCCTGTTTT	ACTTTGTCAT	CAGTTGGATG	720
20	ATTCTCACCT	TTACTCCTCA	GTAAATCAGG	AATGGGAAAT	TAAAAACCAG	TGAATTGAAA	780
	GCACATCTGA	AAGATGCAAT	TCACCATGGA	GCTTTGTCTC	TGGCCCTTAT	TTGTCTAATT	840
25	TTGGAGGTAT	TTGATAACTG	AGTAGGTGAG	GAGATTAAAA	GGGAGCCATA	TAGCACTGTC	900
	ACCCCTTATT	TGAGGAACTG	ATGTTTGAAA	GGCTGTTCTT	TTCTCTCTTA	ATGTCATTTC	960
30	TTTAAAAATA	CATGTGCATA	CTACACACAG	TATATAATGC	CTCCTTAAGG	CATGATGGAG	1020
	TCACCGTGGT	CCATTTGGGT	GACAACCAGT	GACTTGGGAA	GCACATAGAT	ACATCTTACA	1080
	AGTTGAATAG	AGTTGATAAC	TATTTTCAGT	TTTGAGAATA	CCAGTTCAGG	TGCAGCTCTT	1140
35	AAACACATTG	CCTTATGACT	ATTAGAATAT	GCCTCTCTTT	TCATAAATAA	AAATACATGG	1200
	TCTATATCCA	TTTTCTTTTA	TTTCTCTCTC	TTAAGCTTAA	AAAGGCAATG	AGAGAGGTTA	1260
40	GGAGTGGGTT	CATACACGGA	GAATGAGAAA	ACATGCATTA	ACCAATATTC	AGATTTTGAT	1320
	CAGGGGAAAT	TCTAYACTTG	TTGCAAAAAA	AAAAAAAA	AAACTCGAGG	GGGCCCGGT	1380
	ACCCAATCGC	NGTATATGAT	CGNAAACAAT	С			1411
45							
	(2) INFORM	ATION FOR S	EO ID NO: 9	3 :			
50		SEQUENCE C	_				
	ν_,	(A) LEN	GTH: 2187 b	ase pairs			
		(C) STR	ANDEDNESS: OLOGY: line	double			
55	(xi) SEQUENCE			. 93.		
		_			CGTTTCCAAC	TTTCCAGAAG	60
60					TAGATCCCGG		120

	GCGGGCTAAG	AGTAGAATCG	TGTCGCGCTC	GAGAGCGAGA	GTCACGTCCC	GGCGCTAGCC	180
5	CAGCCCGACC	CAGGCCCACC	GTGGTGCACG	CAAACCACTT	CCTGGCCATG	CGCTCCCTCC	240
3	TGCTTCTCAG	CGCCTTCTGC	CTCCTGGAGG	CGGCCCTGGC	CGCCGAGGTG	AAGAAACCTG	300
	CAGCCGCAGC	AGCTCCTGGC	ACTGCGGAGA	AGTTGAGCCC	CAAGGCGGCC	ACGCTTGCCG	360
10	AGCGCAGCCG	GCCTGGCCTT	CAGCTTGTAC	CAGGCCATGG	CCAAGGACCA	GGCAGTGGAG	420
	AACATCCTGG	TGTCACCCGT	GGTGGTGGCC	TCGTCGCTGG	GGCTCGTGTC	GCTGGGCGGC	480
15	AAGGCGACCA	CGGCGTCGCA	GGCCAAGGCA	GTGCTGAGCG	CCGAGCAGCT	GCGCGACGAG	540
	GAGGTGCACG	CCGGCCTGGG	CGAGCTGCTG	CGCTCACTCA	GCAACTCCAC	GGCGCGCAAC	600
	GTGACCTGGA	AGCTGGGCAG	CCGACTGTAC	GGACCCAGCT	CAGTGAGCTT	CGCTGATGAC	660
20	TTCGTGCGCA	GCAGCAAGCA	GCACTACAAC	TGCGAGCACT	CCAAGATCAA	CTTCCGCGAC	720
	AAGCGCAGČG	CGCTGCAGTC	CATCAACGAG	TGGGCCGCGC	AGACCACCGA	CGGCAAGCTG	780
25	CCCGAGGTCA	CCAAGGACGT	GGAGCGCACG	GACGGCGCCC	TGTTAGTCAA	CGCCATGTTC	840
	TTCAAGCCAC	ACTGGGATGA	GAAATTCCAC	CACAAGATGG	TGGACAACCG	TGGCTTCATG	900
	GTGACTCGGT	CCTATACCGT	GGGTGTCATG	ATGATGCACC	GGACAGGCCT	CTACAACTAC	960
30	TACGACGACG	AGAAGGAAAA	GCTGCAAATC	GTGGAGATGC	CCCTGGCCCA	CAAGCTCTCC	1020
	AGCCTCATCA	TCCTCATGCC	CCATCACGTG	GAGCCTCTCG	AGCGCCTTGA	AAAGCTGCTA	1080
35	ACCAAAGAGC	AGCTGAAGAT	CTGGATGGGG	AAGATGCAGA	AGAAGGCTGT	TGCCATCTCC	1140
	TTGCCCAAGG	GTGTGGTGGA	GGTGACCCAT	GACCTGCAGA	AACACCTGGC	TGGGCTGGGC	· 1200
	CTGACTGAGG	CCATTGACAA	GAACAAGGCC	GACTTGTCAC	GCATGTCAGG	CAAGAAGGAC	1260
40	CTGTACCTGG	CCAGCGTGTT	CCACGCCACC	GCCTTTGAGT	TGGACACAGA	TGGCAACCCT	1320
	TTGACCAGAA	TTACGGGCGG	AGGAGTGCGC	ACCCAAGTGT	TCTACGCCGA	CCACCCCTTC	1380
45	ATTTCCTAGT	GCGGGACACC	CAAAGCGGTC	CCTGCTATTC	ATTGGGCGCC	TGGTCCGGCC	1440
	TAAGGGTGAC	AAGATGCGAG	ACGAGTTATA	GGCCTCAGGG	TGCACACAGG	ATGGCAGGAG	1500
	GCATCCAAAG	GCTCCTGAGA	CACATGGGTG	CTATTGGGGT	TGGGGGGGAG	GTGAGGTACC	1560
50	AGCCTTGGAT	ACTCCATGGG	GTGGGGTGGA	AAAGCAGACC	GGGGTTCCCG	TGTGCCTGAG	1620
	CGGACTTCCC	AGCTAGAATT	CACTCCACTT	GGACATGGGC	CCCAGATACC	ATGATGCTGA	1680
 55	GCCCGGAAAC	TCCACATCCT	GTGGGACCTG	GGCCATAGTC	ATTCTGCCTG	CCCTGAAAGT	1740
- -	CCCAGATCAA	GCCTGCCTCA	ATCAGTATTC	ATATTTATAG	CCAGGTACCT	TCTCACCTGT	1800
	GAGACCAAAT	TGAGCTAGGG	GGGTCAGCCA	GCCCTCTTCT	GACACTAAAA	CACCTCAGCT	1860
60	GCCTCCCCAG	CTCTATCCCA	ACCTCTCCCA	ACTATAAAAC	TAGGTGCTGC	AGCCCCTGGG	1920

	ACCAGGCACC CCCAGAATGA CCTGGCCGCA GTGAGGCGGA TTGAGAAGGA GCTCCCAGGA	1980
	HOW DOUGHE CECAMATON CETOGEGOEN GTONOGEGON TITANGANDAN GETECCAGAN	1700
5	GGGGCTTCTG GGCAGACTCT GGTCAAGAAG CATCGTGTCT GGCGTTGTGG GGATGAACTT	2040
_	TTTGTTTTGT TTCTTCCTTT TTTAGTTCTT CAAAGATAGG GAGGGAAGGG GGAACATGAG	2100
	CCTTTGTTGC TATCAATCCA AGAACTTATT TGTACATTTT TTTTTTCAAT AAAACTTTTC	2160
10	CAATGACAAA AAAAAAAAA AAAAAAA	2187
	·	
15	(2) INFORMATION FOR SEQ ID NO: 94:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 757 base pairs (B) TYPE: nucleic acid	
20	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 94:	
25	GACAGTACGG TCGGATTCCC GGGTCGACCC ACGCGTCCGC GGACGGTGAA GAAGGTGAAG	60
	ATGGCGGTGG CCAGGGCCGG GGTCTTGGGA GTCCAGTGGC TGCAAAGGGC ATCCCGGAAC	120
30	GTGATGCCGC TGGGCGCACG GACAGCCTCC CACATGACCA AGGACATGTT CCCGGGGCCC	180
	TATCCTAGGA CCCCAGAAGA ACGGGCCGCC GCCGCCAAGA AGTATAATAT GCGTGTGGAA	240
	GACTACGAAC CTTACCCGGA TGATGGCATG GGGTATGGCG ACTACCCGAA GCTCCCTGAC	300
35 ,	. CGCTCACAGC ATGAGAGAGA TCCATGGTAT AGCTGGGACC AGCCGGGCCT GAGGTTGAAC	360
	TGGGGTGAAC CGATGCACTG GCACCTAGAC ATGTACAACA GGAACCGTGT GGATACATCC	420
40	CCCACACCTG TTTCTTGGCA TGTCATGTGT ATGCAGCTCT TCGGTTTCCT GGCTTTCATG	480
40	ATATTCATGT GCTGGGTGGG GGACGTGTAC CCTGTCTACC AGCCTGTGGG ACCAAAGCAG	540
	TATCCTTACA ATAATCTGTA CCTGGAACGA GGCGGTGATC CCTCCAAAGA ACCAGAGCGG	600
45	GTGGTTCACT ATGAGATCTG AGGAGGCTTC GTGGGCTTTT GGGTCCTCTA ACTAGGACTC	660
	CCTCATTCCT AGAAATTTAA CCTTAATGAA ATCCCTAATA AAACTCAGTG CTGTGTTAAA	720
د ٥	AAAAAAAA AAAAAAAAA AAAAAGGGGG GCCCCNN	757
50		
55	(2) INFORMATION FOR SEQ ID NO: 95:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2394 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 95:

5	GGCACGAGCA	CTCCTGCACT	TCCCCACCCC	CACGACCGAA	CCTGGCTTCG	CTAACGCCCT	60
5	CCCAGCTCCC	TCGGGCCTGA	CTTCCGGTTT	CCTCGCGCGT	CCCTGGCGCC	GAGCCGCGGA	120
	CAGCAGCCCC	TTTTCCGGCT	GAGAGCTCAT	CCACACTTCC	AATCACTTTC	CGGAGTGCTT	180
10	CCCCTCCCTC	CGGCCCGTGC	TGGTCCCGAC	GGCGGGCCTG	GGTCTCGCGC	GCGTATTGCT	240
	GGGTAACGGG	CCTTCTCYCG	CGTCGGCCCG	GCCCTCCTG	CCTCGGCTCG	TCCCTCCTTC	300
15	CAGAACGTCC	CGGCTCCTG	CCGAGTCAGA	AGAAATGGGA	CTCCCTCCGC	GACGTGCCCG	360
13	GAGCAGCTCC	CTTCGCTGTG	GAAGCGGCGG	TGTCTTCGAA	GAAACCGGAA	GCCCGTGGTG	420
	ACCCCTGGCG	ACCCGGTTTG	TTTTCGGTCC	GTTTCCAAAC	ACTAAGGAAT	CGAAACTCGG	480
20	CGGCCTTGGG	GGCGGCCCTA	CGTAGCCTGG	CTTCTGGTTG	TCATGGATGC	ACTGGTAGAA	540
	GATGATATCT	GTATTCTGAA	TCATGAAAAA	GCCCATAAGA	GAGATACAGT	GACTCCAGTT	600
25	TCAATATATT	CAGGAGATGA	ATCTGTTGCT	TCCCATTTTG	CTCTTGTCAC	TGCATATGAA	660
23	GACATCAAAA	AACGACTTAA	GGATTCAGAG	AAAGAGAACT	CTTTGTTAAA	GAAGAGAATA	720
	AGATTTTTGG	AAGAAAAGCT	AATAGCTCGA	TTTGAAGAAG	AAACAAGTTC	CGTGGGACGA	780
30	GAACAAGTAA	ATAAGGCCTA	TCATGCATAT	CGAGAGGTTT	GCATTGATAG	AGATAATTTG	840
	AAGAGCAAAC	TGGACAAAAT	GAATAAAGAC	AACTCTGAAT	CTTTGAAAGT	ATTGAATGAG	900
35	CAGCTACAAT	CTAAAGAAGT	AGAACTCCTC	CAGCTGAGGA	CAGAGGTGGA	AACTCAGCAG	960
33	GTGATGAGGA	ATTTAAATCC	ACCTTCATCA	AACTGGGAGG	TGGAAAAGTT	GAGCTGTGAC	1020
	CTGAAGATCC	ATGGTTTGGA	ACAAGAGCTG	GAACTGATGA	GGAAAGAATG	TAGCGATCTC	1080
40	AAAATAGAAC	TACAGAAAGC	CAAACAAACG	GATCCATATC	AGGAAGACAA	TCTGAAGAGC	1140
	AGAGATCTCC	AAAAACTAAG	CATTTCAAGT	GATÄATATGC	AGCATGCATA	CTGGGAACTG	1200
45	AAGAGAGAAA	TGTCTAATTT	ACATCTGGTG	ACTCAAGTAC	AAGCTGAACT	ACTAAGAAAA	1260
43	CTGAAAACCT	CAACTGCAAT	CAAGAAAGCC	TGTGCCCCTG	TAGGATGCAG	TGAAGACCTT	1320
	GGAAGAGACA	GCACAAAACT	GCACTTGATG	AATTTTACTG	CAACATACAC	AAGACATCCC	1380
50	CCTCTCTTAC	CAAATGGCAA	AGCTCTTTGT	CATACCACAT	CTTCCCCTTI	ACCAGGAGAT	1440
	GTAAAGGTTT	TATCAGAGAA	AGCAATCCTC	CAATCATGGA	CAGACAATGA	GAGATCCATT	1500
55	CCTAATGATG	GTACATGCTT	TCAGGAACAC	AGTTCTTATO	GCAGAAATTC	TCTGGAAGAC	1560
טט	AAT¶CCTGGG	TATTTCCAAG	TCCTCCTAAA	TCAAGTGAGA	CAGCATTTGC	GGAAACTAAA	1620
	ACTAAAACTT	TGCCTTTACC	CAACCTTCCA	CCACTGCATT	ACTTGGATC	A ACATAATCAG	1680
60	AACTGCCTTT	ATAAGAATTA	ATTTGGAAGA	GATTCACGAT	TTCACCATG	A GGACACTTAT	1740

480

540 ..

600

660

672

	CTCTTTCAGT GG	TCCTCCCA .	AGAAATTATT	TAACAAACTG	AANGGAGATT	TTGATTAAAA	1800
5	TTTTGCAGAG GT	CTTCAGTA	TCTATATTTG	AACACACTGT	ACAATAGTAC	AAAAACCAAC	1860
J	ATAGTTGGTT TT	CTAGTATO	AAAGAGCACC	CTCTAGCTCC	ATATTCTAAG	AATCTGAAAT	1920
	ATGCTACTAT AC	TAATTAAT	AAGTAAACTT	AAGGTGTTTA	AAAAACTCTG	ССТТСТАТАТ	1980
10	TAATTGTAAA AT	TTTGCCTC	TCAGAAGAAT	GGAATTGGAG	ATTGTAGACG	TGGTTTTACA	2040
	AAATGTGAAA TG	TCTAAATA	TCTGTTCATA	AAAATAAAAG	GAAAACATGT	TTCTTCAAAT	2100
15	TGCATAATGG AA	CAAATGGC .	aatgtgagta	GGTTACATTT	CTGTTGTTAT	AATGCGTAAA	2160
13	GATATTGAAA AT	TATAATGAA	ATAAAAGCAT	CTTAGGTTAT	ACCATCTTTA	TATGCTATTG	2220
	CGTTTCAATA TT	TAAGATTT	AAAGTGATTT	TTTGGTCACA	GTGTTTTGTT	GATAAAATTT	2280
20	TTTTAGAATT GA	AGTTTGAA	TTCTAAGACT	TGAAACAACC	TGATCACTGA	AGCCAACTTT	2340
	GTCCCAGCAC AT	TCCTTAAG	TCCTAATTGG	GGAAAAAAA	АААААААА	TCGA	2394
25							
20	(2) INFORMATI	ON FOR SE	O TO NO. 94				
	(2) INFORMATI	ION FOR SE	Q ID NO: 36				
30	(i) SI	(A) LENC (B) TYPE (C) STRA	WARACTERIST GTH: 672 ba E: nucleic ANDEDNESS: DLOGY: line	se pairs acid double			
35	(xi) 5	SEQUENCE I	ESCRIPTION	: SEQ ID NC	: 96:		
	AGTGCTCTGT TO	GCCAGGCT	GGAGTGCGTT	AGTGTAATGT	CAGTCCACTG	CAACCTCCAC	60
40	CCCCAGGTTC A	AGCAATTCT	CATGCCTCAG	CCTCCCAAGT	AGCTGAAATT	ACTGGCATGC	120
40	ACCACCACAC CO	CAGCTGATG	TTTATTTATT	ТАТТТАТАТА	TTTATTTATT	TTAGGTGTTT	180
	TTTTTTTTT T	ITTTGAGAC	GGAGTCTTGC	TCTGTTGCCC	TGGGTGTGGT	TACGTGGRAT	240
45	TACCATYCTG GO	GTGACTCAC	TGAAATGTAC	TCMCAGTGAG	TCATGCCTTC	MAATGACATC	300
	TCAAGTTCTG CO	CTGCTTGGA	GATACATCTG	GGGATCTTAA	GGGGTGAGGG	ACTACTCAAC	360

AAGAAGGAAT TTAGCCTGTC TTTTTAAATA AACGGCATTT CTTTTTCCTA KAAAAATGGG

AAATTCTTCA ATTCTCTAAT ACAGGACAC TGAGATAACA AAGAGGAAAG TGTCTGGTTG

AAATCMAGCA AAGCACAARA AAKTTTCCCT TTGCTAAAAG GGAAAAGATG CCCCMCAATG

CCCATAAACA TGAACTGGGG ATAAGGAGGA RAATGTCTCT YCTTGGCACC CCCAAACAAA

60

CGTTAATTAC CC

50

WO 98/42738 PCT/US98/05311

248

(2) INFORMATION FOR SEQ ID NO: 97	(2)	INFORMATION	FOR	SEO	ID	NO:	97
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(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1419 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

10 (D) TOPOLOGY: linear

; (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 97:

	•	_					
15	TAAGAACAGA	ACAGCAAGTA	TGAACCACAT	GGAACTTAAA	ACATATGGGT	GTGAAGTCCA	60
	CTTATGTAGA	CAAAACTTAT	AATTTCCAAA	CTGTTGTCTA	GTATACAGTG	ATCAGTTGCT	120
	CTCTGTTCAA	GTCATTCCAC	ACATTTCCCT	ATTTTAGGCT	АТТАТААТАТ	AGAAAGAAAA	180
20	TGGGAAGCAT	TAGTTGGAGC	TAGAAAATGA	ACTGTATATT	ATTGCTATAT	TTGCTAATAC	240
	CAACTATTTC	AATAAGTGTT	GTACCATATG	TAGCATTAAA	TATAAAATAC	ATAAAAGAAT	300
25	GTACAGAAAA	TAGCTTTTAT	TGAGTAATAT	TACATTTCAT	TTATACTGTA	GCAATATATT	360
23	TGTAGGTATA	CTCTGTAAGG	GCTTTAAATA	AAAGAGGTCC	ATTAATACTT	CCTTATAAAA	420
	ATTCTAGTCT	GTTTCATTAC	TGCCCAGATG	TTTTAGAGAT	AAATATTTAT	GCAGAAGGTA	480
30	TTTTKGAAAG	TCYCCYTTIG	TCTGATAGAG	TTTAACNAGA	TATTTAAATT	TAGTGCYCNA	540
	GAAATCCCAC	AAGTCACGGT	CTAAACACAC	TTAGAATACT	ACAGCATAAA	TCTGTTAGCA	600
35	TTANTTGCCA	AATAAGACAG	TTGGGATCCC	AAACCCCAAG	TCCTTGAGCA	ATGTTTTTCC	660
33	TCAAAAAGCT	GCTATNCCAA	TGATATAGGA	AAAWACATTG	TGTTTTCCTA	AACACACTTT	720
	TCTTTTTAAA	TGTGCTTCAT	TGTTTGATTT	GGTCCTGCCT	AAATTTCACA	AGCTAGGCCA	780
40	ATGAAGGCTG	AATCAAAGAC	ATTTCATCCA	CCAATATCAT	GTGTAGATAT	TATGTATAGA	840
	ATAAAATAA	. AATTATGGCT	CTAACTTCTG	TGTTGCTGTT	TATCTTGTTA	TTTTTCGGCG	900
45	TTATACTAAT	GNGTTTATTG	AGAGCATTTI	' ACCTTCCAGA	CTTCTCATGO	CTAACTTTTG	960
43	GTCTGWATTT	'TGSTCCTTAG	ATGKGAATAT	TTCTTATTAG	TYTGCTYCCT	GCWACGCAAT	1020
	GACTGCATTT	CTATCATTTC	TCAGTTTGT	AGWATATGTO	GATAGTATTO	TACTGTATAA	1080
50	ATGATTGCAA	AGTTTATCAA	. AAACAAATTA	TTATATGTAG	CTTTTCTACE	A GTGCTTTGCT	1140
	AAACCATGTA	GTACTAGTTA	AGTSTTCCT	GAAAATAAAC	ATACACTCT	r ataggggaca	1200
55	GTTCCTGTTC	ACTCCCAGGA	AACTTTTT	A AAAGATGACA	CTGAATGTT	r ATTGCACTTT	1260
33	AGTGCAGTGA	AGTGGCAATA	AAACCTAAC	A TGAATCAAGO	TTGTTTATG	G CAGATGCATG	1320
	TGTTGCTTTA	A CAGAGTTTAC	CAAAAGCTC	TAATTTATO	G TCATACTGT	A TTCTACTGAA	1380
60	TAATAAAGCT	T AACATTATTO	AATAATAA	A TGGAAAAA			1419

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`	121	INFORMATION	EUB	SEO	TD	MO .	QQ.

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1487 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

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	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 98:	
15	GCGACCGCGC CCCTTTCAGC TAGCTCGCTC GCTCGCTCTG CTTCCCTGCT GCCGGCTGCG	60
	CATGGCKWTG GCGTTGGCGG CGCTGGCGGC GGTCGAGCCG GCCTGCGCAG CCGGTACCAG	120
20	CAGTTGCAGA ATGAAGAAGA GTCTGGAGAA CCTGAACAGG CTGCAGGTGA TGCTCCTCCA	180
20	CCTTACAGCA GCATTTCTGC AGAGAGCGCA GTTTTCCACC TATTTCCCTG GATATTTTGA	240
	TGGTCAGTAC TGGCTCTGGT GGGTGTTCCT TGTTTTAGGC TTTCTCCTGT TTCTCAGAGG	300
25	ATTTATCAAT TATGCAAAAG TTCGGAAGAT GCCAGAAACT TTCTCAAATC TCCCCAGGAC	360
	CAGAGTTCTC TTTATTTATT AAAGATGTTT TCTGGCAAAG GCCTTCCTGC ATTTATGAAT	420
30	TCTCTCTCAA GAAGCAAGAG AACACCTGCA GGAAGTGAAT CAAGATGCAG AACACAGAGG	480
	AATAATCACC TGCTTTAAAA AAATAAAGTA CTGTTGAAAA GATCATTTCT CTCTATTTGT	540
	TCCTAGGTGT AAAATTTTAA TAGTTAATGC AGAATTCTGT AATCATTGAA TCATTAGTGG	600
35	TTAATGTTTG AAAAAGCTCT TGCAATCAAG TCTGTGATGT ATTAATAATG CCTTATATAT	660
	TGTTTGTAGT CATTTTAAGT AGCATGAGCC ATGTCCCTGT AGTCGGTAGG GGGCAGTCTT	720
40	GCTTTATTCA TCCTCCATCT CAAAATGAAC TTGGAATTAA ATATTGTAAG ATATGTATAA	780
	TGCTGGCCAT TTTAAAGGGG TTTTCTCAAA AGTTAAACTT TTGTTATGAC TGTGTTTTTG	840
	CACATAATCC ATATTTGCTG TTCAAGTTAA TCTAGAAATT TATTCAATTC TGTATGAACA	900
45	CCTGGAAGCA AAATCATAGT GCAAAAATAC ATTTAAGGTG TGGTCAAAAA TAAGTCTTTA	960
	ATTGGTAAAT AATAAGCATT AATTTTTAT AGCCTGTATT CACAATTCTG CGGTACCTTA	1020
50	TTGTACCTAA GGGATTCTAA AGGTGTTGTC ACTGTATAAA ACAGAAAGCA CTAGGATACA	1080
	AATGAAGCTT AATTACTAAA ATGTAATTCT TGACACTCTT TCTATAATTA GCGTTCTTCA	1140
	CCCCCACCC CACCCCCACC CCCCTTATTT TCCTTTTGTC TCCTGGTGAT TAGGCCAAAG	1200
55	TCTGGGAGTA AGGAGGGAT TAGGTACTTA GGAGCAAAGA AAGAAGTAGC TTGGAACTTT	1260
	TGAGATGATC CCTAACATAC TGTACTACTT GCTTTTACAA TGTGTTAGCA GAAACCAGTG	1320
60	GGTTATAATG TAGAATGATG TGCTTTCTGC CCAAGTGGTA ATTCATCTTG GTTTGCTATG	1380

1320

250

	TTAAAACTGT AAATACAA	ACA GAACATTAAT	AAATATCTCT	TGTGTAGCAC	CTTTAAAAAA	1440
	KAAAAAA AAAAAAAA	NAAAAAAAA AA	CCCGGGGGGG	GGCCCCN		1487
5						
_	(2) INFORMATION FOR	R SEQ ID NO: 99	9:			
.0		E CHARACTERIST: LENGTH: 1653 b				
	• •	TYPE: nucleic STRANDEDNESS:				
.5	(D)	TOPOLOGY: line	ar			
	(xi) SEQUEN	CE DESCRIPTION	: SEQ ID NO	: 99:		
	GCGACCGCGC CCTTCAGC	CTA GCTCGCTCGC	TCGCTCTGCT	TCCCTGCTGC	CGGCTGCGCA	60
20	TGGCTTNGGC GTTGGCGC	SCG CTGGCGGCGG	CTCGAGCCGC	CTGCGSAGCC	GGTACCAGCA	120
	GTTGCAGAAT GAAGAAGA	AGT CTGGAGAACC	TGAACAGGCT	GCAGGTGATG	CTCCTCCACC	180
25	TTACAGCAGC ATTTCTGG	CAG AGAGCGCACA	TNATTTTGAC	TACAAGGATG	AGTCTGGGTT	240
	TCCAAAGCCC CCATCTTA	ACA ATGTAGCTAC	AACACTGCCC	AGTTATGATG	AAGCGGAGAG	300
	GACCAAGGCT GAAGCTAG	CTA TCCCTTTGGT	TCCTGGGAGA	GATGAGGATT	TTGTGGGTCG	360
80	GGATGATTTT GATGATGO	CTG ACCAGCTGAG	GATAGGAAAT	GATGGGATTT	TCATGTTAAC	420
	TTTTTCATG GCATTCC	TCT TTAACTGGAT	TGGGTTTTTC	CTGTCTTTTT	GCCTGACCAC	480
35	TTCAGCTCCA GGAAGGT	ATG GGGCCATTTC	AGGATTTGGT	CTCTCTCTAA	TTAAATGGAT	540
, ,	CCTGATTGTC AGGTTTTC	CCA CCTATTTCCC	TGCATTTATG	AATTCTCTCT	CAAGAAGCAA	600
	GAGAACACCT GCAGGAAG	GTG AATCAAGATG	CAGAACACAG	AGGAATAATC	ACCTGCTTTA	660
10	AAAAAATAAA GTACTGT	IGA AAAGATCATT	TCTCTCTATT	TGTTCCTAGG	TGTAAAATTT	720
	TAATAGTTAA TGCAGAA	ITC TGTAATCATT	GAATCATTAG	TGGTTAATGT	TTGAAAAAGC	780
15	TCTTGCAATC AAGTCTG	IGA TGTATTAATA	ATGCCTTATA	TATTGTTTGT	AGTCATTTTA	840
+3	AGTAGCATGA GCCATGT	CCC TGTAGTCGGT	AGGGGGCAGT	CTTGCTTTAT	TCATCCTCCA	900
	TCTCAAAATG AACTTGG	AAT TAAATATTGT	AAGATATGTA	TAATGCTGGC	CATTTTAAAG	960
50	GGGTTTTCTC AAAAGTT	AAA CTTTTGTTAT	GACTGTGTTT	TTGCACATAA	TCCATATTTG	1020
	CTGTTCAAGT TAATCTA	GAA ATTTATTCAA	TTCTGTATGA	ACACCTGGAA	GCAAAATCAT	1080
	AGTGCAAAAA TACATTT	AAG GTGTGGTCAA	AAATAAGTCT	TTAATTGGTA	 AATAATAAGC	1140
55	ATTAATTTT TATAGCC	TGT ATTCACAATT	CTGCGGTACC	TTATTGTACC	TAAGGGATTC	1200
	TAAAGGTGTT GTCACTG	TAT AAAACAGAAA	GCACTAGGAT	' ACAAATGAAG	CTTAATTACT	1260

AAAATGTAAT TCTTGACACT CTTTCTATAA TTAGCGTTCT TCACCCCCAC CCCCACCCCC

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	ACCCCCCTTA TTTTCCTTTT GTCTCCTGGT GATTAGGCCA AAGTCTGGGA GTAAGGAGAG	1380
5	GATTAGGTAC TTAGGAGCAA AGAAAGAAGT AGCTTGGAAC TTTTGAGATG ATCCCTAACA	1440
3	TACTGTACTA CTTGCTTTTA CAATGTGTTA GCAGAAACCA GTGGGTTATA ATGTAGAATG	1500
	ATGTGCTTTC TGCCCAAGTG GTAATTCATC TTGGTTTGCT ATGTTAAAAC TGTAAATACA	1560
10	ACAGAACATT AATAAATATC TCTTGTGTAG CACCTTTTAW AAAAAAAAA AAAAAAAAA	1620
	AAAAAAAA AAAAANCCCG GGGGGGGCC CCN	1653
15		
	(2) INFORMATION FOR SEQ ID NO: 100:	
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1145 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 100:	
	TITTTTTTT TTTTTTTTT TTGACTGAAC TAAGTGGCTT TTTTATTAGA GAAAGCCAGA	60
30	ATTACAAAAG ACTTCCCTTT TCTTGGGGTA TGGCTGTCTC AGCACAATAC TCAACATAAC	120
,,,	TGCAGAACTG ATGTGGCTCA GGCACCCTGG TTTTAATTCC TTGAGGATCT GGCAATTGGC	180
	TTACGCAAAA GGTCACCATT TGAGGTCCTG CCTTACTAAT TATGTGCTGC CCAACAACTA	240
35	AATTTGTAAT TTGTTTTTCT CTAGTTTGAG CAGGGTCTGA ATTTTTTCCTT	300
	TTTTGCCAGC AGACAGACTT GAGTCTGTAA AGACAAGCAA ATACACTGAC AGAAGTTTAC	360
40	CATAGTTTCT AAAATGTAAA AAAGAAAACC CCCAAAAGAC TCAAGAAAAT TAGACCACAA	420
	ATTITICATT GTTCATTGTA GCACTATTGG TAATAAAATA ACAAATGTTT GTGCATTTTT	480
	ATGTGAAGAT CCTTCTCGTA TTTCATTTGG AAAGATGAGC AAGAGGTCTG CTTCCTTCAT	540
45	TTTACTTCCC CTTCTGTTTT TGAAAGGCAG TTTCGCCAAG CTTAATGCAA GAATATCTGA	600
	CTGTTTAGAA GAAAGATATT GCCACAATCT CTGGATGGTT TTCCAGGGTT GTGTTATTAC	660
50	TGAGCTTCAT CTTTCCAGAA TGAGCAAAAC ACTGTCCAGT CTTTGTTACG ATTTTGTAAT	720
	AAATGTGTAC ATTTTTTTTA AATTTTTGGA CATCACATGA ATAAAGGTAT GTATGTACGA	780
	ATGTGTATAT ATTATATATA TGACATCTAT TTTGGAAAAT GTTTGCCCTG CTGTACCTCA	84
55	TTTTTAGGAG GTGTGCATGG ATGCAATATA TGAAAATGGG ACATTCTGGA ACTGCTGGTC	90
	AGGGGACTTT GTCGCCCTGT GCACTAAAAG GGCCAGATTT TCAGCAGCCA AGGACATCCA	96
	TACCCAAGTG AATGTGATGG GACTTAAAAG AAGTGAACTG AGACAATTCA CTCTGGCTGT	102

	TTGAACAGCA GCGTTTCATA GGAAGAGAAA AAAAGATCAA TCTTGTATTT TCTGACCACA	1080
	TAAAGGCTTC TTCTCTTTGT AATAAAGTAG AAAAGCTCTC CTCAAAAAAA AAAAAAAAAA	1140
5	AAAAA	1145
	•	
10	(2) INFORMATION FOR SEQ ID NO: 101:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 734 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 101:	
20	TACCCGGCGG ATTCCAGGAA GGTAAATTTA GTCCTATAAT TTTCAGCTTA ATTATAAACA	60
	AAGGAACAAA TAAGTGGAAG GGCAGCTATT ACCATTCGCT TAGTCAAAAC ATTCGGTTAC	120
25	TGCCCTTTAA TACACTCCTA TCATCAGCAC TTCCACCATG TATTACAAGT CTTGACCCAT	180
	CCCTGTCGTA ACTCCAGTAA AAGTTACTGT TACTAGAAAA TTTTTATCAA TTAACTGACA	240
	AATAGTTTCT TTTTAAAGTA GTTTCTTCCA TCTTTATTCT GACTAGCTTC CAAAATGTGT	300
30	TCCCTTTTTG AATCGAGGTT TTTTTGTTTT GTTTTGTTTT	360
	TGTGCTTCTA TTGCTTTTTT GTGTTTTGTT AAGCATGTCC CTTGGCCCAA ATGGAAGAGG	420
35	AAATGTTTAA TTAATGCTTT TTAGTTTAAA TAAATTGAAT CATTTATAAT AATCAGTGTT	480
	AACAATTTAG TGACCCTTGG TAGGTTAAAG GTTGCATTAT TTATACTTGA GATTTTTTTC	540
	CCCTAACTAT TCTGTTTTTT GTACTTTAAA ACTATGGGGG AAATATCACT GGTCTGTCAA	600
40	GAAACAGCAG TAATTATTAC TGAGTTAAAT TGAAAAGTCC AGTGGACCAG CCATTTCTTA	660
	TATAAATAAA ATTGGTGGTA CTAATGTGAA AAAAAAAAAA	720
45	CCGGTACCCT ATTA	734
50	(2) INFORMATION FOR SEQ ID NO: 102:	
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 713 base pairs(B) TYPE: nucleic acid	
55	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 102:	
60	CCGCGGGAAC GCTGTCCTGG CTGCCGNCAC CCGAACAGCC TGTCCTGGTG CCCCGGCTCC	60

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	CTGCCCCGCG	CCCAGTCATG	ACCCTGCGCC	CCTCACTCCT	CCCGCTCCAT	CTGCTGCTGC	120
	TGCTGCTGCT	CAGTGCGGCG	GTGTGCCGGG	CTGAGGCTGG	GCTCGAAACC	GAAAGTCCCG	180
5	TCCGGACCCT	CCAAGTGGAG	ACCCTGGTGG	AGCCCCCAGA	ACCATGTGCC	GAGCCCGCTG	240
	CTTTTGGAGA	CACGCTTCAC	ATACACTACA	CGGGAAGCTT	GGTAGATGGA	CGTATTATTG	300
10	ACACCTCCCT	GACCAGAGAC	CCTCTGGTTA	TAGAACTTGG	CCAAAAGCAG	GTGATTCCAG	360
10	GTCTGGAGCA	GAGTCTTCTC	GACATGTGTG	TGGGAGAGAA	GCGAAGGGCA	ATCATTCCTT	420
	CTCACTTGGC	CTATGGAAAA	CGGGGATTTC	CACCATCTGT	CCCAGCGGAT	GCAGTGGTGC	480
15	AGTATGACGT	GGAGCTGATT	GCACTAATCC	GAGCCAACTA	CTGGCTAAAG	CTGGTGAAGG	540
	GCATTITGCC	TCTGGTAGGG	ATGGCCATGG	TGCCACCCTC	CTGGGCCTCA	TTGGGTATCA	600
20	CCTATACAGA	AAGGCCAATA	GACCCAAAGT	CTCCAAAAAG	AAGCTCAAGG	AAGAGAAACG	660
	AAACAAGAGC	AAAAAGAAAT	AATAAATAAT	AAATTTTAAA	AAACTTAAAA	AAA	713
25	(2) INFORM	ATION FOR S	EQ ID NO: 1	03:			

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1080 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 103:

35 CCGATGTGGA CATCATCCTG TCTATCCCCA TGTTCCTGCG CCTGTACCTG ATCGCCCGAG 60 TCATGCTGCT GCACAGAAGC TCTTCACCGA TGCCTCGTCC CGCAGCATCG GGGCCCTCAA 120 40 CAAGATCAAC TTCAACACCC GCTTTGTCAT GAAGACGCTC ATGACCATCT GCCCTGGCAC 180 TGTGCTGCTC GTGTTCAGCA TCTCTCTGTG GATCATTGCT GCCTGGACCG TCCGTGTCTG 240 TGAAAGTCCT GAATCACCAG CCCAGCCTTC TGGCTCATCA CTTCCTGCTT GGTACCATGA 300 45 CCAGCAGGAC GTAACTAGTA ACTITCTGGG TGCCATGTGG CTCATCTCCA TCACATTCCT 360 TTCCATTGGT TATGGGGACA TGGTGCCCCA CACATACTGT GGGAAAGGTG TCTGTCTCCT 420 50 CACTGGCATC ATGGGTGCAG GCTGCACTGC CCTTGTGGTG GCCGTGGTGG CCCGAAAGCT 480 GGAACTCACC AAAGCGGAGA AGCACGTTCA TAANTTCATG ATGGACACTC AGCTCACCAA 540 GCGGATCAAG AATGYTGCAG CCAATGTCCT TSGGGAAACA TGGTTAATCT ATAAACACAC 600 55 AAAGYTGYTA AAGAAGATTG ACCATGCCAA AGTGAGGAAC ACCAGAGGAA GTTCYTCCAA 660 GTATCCACCA GTTGAGGAGC GTCAAGATGG AACAGAGGAA GCTGAGTGAC CAAGCCAACA 60 NTCTGGTGGA CCTTTCCAAG ATGCAGAATG TCMTGTATGA CTTAATCACA GAACTCAATG

	ACCGGAGCGA AGACCTGGAG AAGCAGATTG GCAGCCTGGA GTCGAAGCTG GAGCATCTCA	840
5	CCGCCAGCTT CAACTCCCTG CCGCTGCTCA TCGCCGACAC CCTGCGCCAG CAGCAGCAGC	900
)	AGCTCCTGTC TGCCATCATC GAGGCCCGGG GTGTCAGCGT GGCAGTGGGC ACCACCCACA	960
	CCCCAATCTC CGATAGCCCC ATTGGGGTCA GCTCCACCTC CTTCCCGACC CCGTACACAA	1020
10	GTTCAAGCAG TTGCTAAATA AATCTCCCCA CTCCAGAAGC ATTAAAAAAA AAAAAAAAA	1080
15	(2) INFORMATION FOR SEQ ID NO: 104:	
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 489 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 104:	
25	GGCACGAGAG GCTTTGAAGC ATTTTTGTCT GTGCTCCCTG ATCTTCAGGT CACCACCATG	60
	AAGTTCTTAG CAGTCCTGGT ACTCTTGGGA GTTTCCATCT TICTGGTCTC TGCCCAGAAT	120
30	CCGACAACAG CTGCTCCAGC TGACACGTAT CCAGCTACTG GTCCTGCTGA TGATGAAGCC	180
50	CCTGATGCTG AAACCACTGC TGCTGCAACC ACTGCGACCA CTGCTGCTCC TACCACTGCA	240
	ACCACCGCTG CTTCTACCAC TGCTCGTAAA GACATTCCAG TTTTACCCAA ATGGGTTGGG	300
35	GATCTCCCGA ATGGTAGAGT GTGTCCCTGA GATGGAATCA GCTTGAGTCT TCTGCAATTG	360
	GTCACAACTA TTCATGCTTC CTGTGATTTC ATCCAACTAC TTACCTTGCC TACGATATCC	420
40	CCTTTATCTC TAATCAGTTT ATTTTCTTTC AAATAAAAAA TAACTATGAG CAACAAAAAA	480
	AAAAAAAA	489
45	(2) INFORMATION FOR SEQ ID NO: 105:	
	(i) SEQUENCE CHARACTERISTICS:	
50	(A) LENGTH: 640 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 105:	
55	GCGGTCGCCG CTGTTGTTGT GGTCCCCATG GAGCTGCCGT AGCGGACCCA GCACAGCCAG	60
	GAGCGTCCGG GATGAGCTCA GCCGCGGCCG ACCACTGGGC GTGGTTGCTG GTGCTCAGCT	120
60	TCGTGTTTGG ATGCAATGTT CTTAGGATCC TCCTCCCGTC CTTCTCATCC TTCATGTCCA	180

780

840

900

	GGGTGCTGCA GAAGGACGCG GAGCAGGAGT CACAGATGAG AGCGGAGATC CAGGACATGA	240
5	AGCAGGAGCT CTCCACAGTC AACATGATGG ACGAGTTTGC CAGATATGCC AGGCTGGAAA	300
3	GAAAGATCAA CAAGATGACG GATAAGCTCA AAACCCATGT GAAAGCTCGG ACAGCTCAAT	360
	TAGCCAAGAT AAAATGGGTG ATAAGTGTCG CTTTCTACGT ATTGCAGGCT GCCCTGATGA	420
10	TCTCACTCAT TTGGAAGTAT TATTCTGTCC CTGTGGCTGT CGTGCCGAGT AAATGGATAA	480
	CCCTYTAGAC CGCCTGGTAG CCTTTCCYAY TAGAGTAGCA GGTGGTGTTG GAATTACTGT	540
	TGGATTTART CTGTACAAAT TGTCCTATTG TGCTTCACCG TYCASTGAAC AGGAGGTGGT	600
15	ACAGCCGGAG TTAAAAAACGG TTTCCNTTCC AGTTTAAAAT	640
20		
	(2) INFORMATION FOR SEQ ID NO: 106:	
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1529 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 106:	
30	GGGCACNAGA TGGAGCTGCC GTAGCGGACC CAGCACAGCC AGGAGCGTCC GGGATGAGCT	60
	CAGCCGCGGC CGACCACTGG GCGTGGTTGC TGGTGCTCAG CTTCGTGTTT GGATGCAATG	120
35	TTCTTAGGAT CCTCCTCCCG TCCTTCTCAT CCTTCATGTC CAGGGTGCTG CAGAAGGACG	180
	CGGACAGGAG TCACAGATGA GAGCGGAGAT CCAGGACATG AAGCAGGAGC TCTCCACAGT	240
40	CAACATGATG GACGAGTTTG CCAGATATGC CAGGCTGGAA AGAAAGATCA ACAAGATGAC	300
40	GGATAAGCTC AAAACCCATG TGAAAGCTCG GACAGCTCAA TTAGCCAAGA TAAAATGGGT	360
	GATAAGTGTC GCTPTCTACG TATTGCAGGC TGCCCTGATG ATCTCACTCA TTTGGAAGTA	420
45	TTATTCTGTC CCTGTGGCTG TCGTGCCGAG TAAATGGATA ACCCCTCTAG ACCGCCTGGT	480
	AGCCTTTCCT ACTAGAGTAG CAGGTGGTGT TGGAATTACC TGTTGGATTT TAGTCTGTAA	540
50	CAAAGTTGTC GCTATTGTGC TTCATCCGTT CAGCTGAACA GGAGGATGGA TACAGCCGCG	600
50	AGTAAAAAA CGGATTTCCT CTTCCTAGCT TAAAATCTGA TTTACACTGT TTTGTTTTTT	660
	AAGAAACAAA AGTGCATAGT TTAGATTTYT TTTTTGTTGA ATATGTTTGT TCTTGGACTT	720

55 TATGAGATAG TCTTATAAGA ATCACGATTT TCTACACCTG TCATTGAGCC AAGAAAGTCC

AGTTTATGAC ACGTATGTAC TAGTGAACAC CGTCCTCGAT CTGTACGAAA TGTGAAATGT

TTAGGGACAT CTCCATGCTG TCACTTGTGA TTTGCCCTCT TATGTATTTT GGTCATATTG

	CCAACTGGAA	AGTCAAAATT	TTCTAACAAC	TTTAAGTAAG	TTCTTTGAAG	ACTTAGTGCT	960
	GTTTTTAATC	CAGTTTAGAA	AGTAACTTAA	TTTTAATACC	RCTACTAAAA	ATTCGAAAAT	1020
5	TTCTTCTTTA	ATCACATTCA	ATATGGTTAA	AAGAACAACA	CTAATTGACA	TTGCGTGGGC	1080
	TTTTTCTCCC	TTTGTTTAAA	ATGTCATTTG	TTGAGCAAGA	GTTGTATAGT	ATTATCTACT	1140
10	TACTTGAGGC	TGTTAATTTT	TCATTACAGT	GTTTTGTAAA	TGTATCCACG	AGACCATGAT	1200
10	GCATTGTTTT	GTGCTCAACT	TGTGTTTTGT	ATTTAAAGCA	TTTTGAATGA	AGTGTATTTT	1260
	ATAAGCATTT	AATATTTATG	CTCTTTAGAA	TGGAACACAG	AAAACAAACC	TTATAAGTCC	1320
15	TGATTAATCT	GAACCAATAA	CCTGTGTGGC	CTACAAAGTA	TAATTCTATT	AAATGTTCCT	1380
	TAAAACACTT	TTTTCTAATT	AAAATCTTTG	CAAATGCTTG	TGTAACTTCC	TGCCTTACAG	1440
20	CTACTTGTTT	GCTGTGAGCC	ACCCGCAACT	GACAAGTGGC	TGTTAACTGA	GTCACCATAT	1500
20	CCCAGTAAAG	CTGAATTTTC	TCACTAAAA				1529

(2) INFORMATION FOR SEQ ID NO: 107:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2435 base pairs

30 (B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 107:

35	(21,	SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 107:		
ŞS	ATGAAGGGTC	GTTGGTGGGA	AAGATGGCGG	CGACTCTGGG	ACCCCTTGGT	CGTGGCAGCA	60.
	GTGGCGRCGA	TGTTTGTCGG	CTCGGGATGG	GTCCAGGATG	TTACTCCTTC	TTCTTTTGTT	120
40	GGGGTCTGGG	CAGGGGCCAC	AGCAAGTCGG	GGCGGGTCAA	ACGTTCGAGT	ACTTGAAACG	180
	GGAGCACTCG	CTGTCGAAGC	CCTACCAGGG	TGTGGGCACA	GGCAGTTCCT	CACTGTGGAA	240
45	TCTGATGGGC	AATGCCATGG	TGATGACCCA	GTATATCCGC	CTTACCCCAG	ATATGCAAAG	300
43	TAAACAGGGT	GCCTTGTGGA	ACCGGGTGCC	ATGTTTCCTG	AGAGACTGGG	AGTTGCAGGT	360
	GCACTTCAAA	ATCCATGGAC	AAGGAAAGAA	GAATCTGCAT	GGGGATGGCT	TGGCAATCTG	420
50	GTACACAAAG	GRWTCGGATG	CAGCCAGGGC	CTGTNTTTGG	GAAACATGGA	CAAATTTGTG	480
	GGGCTGGGAG	TATTTGTAGA	CACCTACCCC	AATGAGGAGA	AGCAGCAAGA	GCGGGTATTC	540
55	CCCTRCMTCT	CAGCCATGGT	GAACAACGGC	TCCCTCAGCT	ATGATCATGA	GCGGGATGGG	600
55	CGGCCTACAG	AGCTGGGAGG	CTGCASAGCC	ATTGTCCGCA	ATCTTCATTA	CGACACCTTC	. 660
	CTGGTGATTC	GCTACGTCAA	GAGGCATTTR	ACGATAATGA	TGGATATTGA	TGGCAAGCAT	720
60	GAGTGGAGGG	ACTGCATTGA	AGTGCCCGGA	GTCCGCCTGC	CCCGCGGCTA	CTACTTCGGC	780

	ACCTCCTCCA	TCACTGGGGA	TCTCTCAGAT	AATCATGATG	TCATTTCCTT	GAAGTTGTTT	840
5	GAACTGACAG	TGGAGAGAAC	CCCAGAAGAG	GAAAAGCTCC	ATCGAGATGT	GTTCTTGCCC	900
	TCAGTGGACA	ATATGAAGCT	GCCTGAGATG	ACAGCTCCAC	TGCCGCCCCT	GAGTGGCCTG	960
	GCCCTCTTCC	TCATCGTCTT	TTTCTCCCTG	GGTGTTTTCT	GTATTTGCCA	TAGTCATTGG	1020
10	TATCATACTC	TACAACAAAT	GGCAGGAACA	GAGCCGAAAG	CGCTTCTACT	GAGCCCTCCT	1080
	GCTGCCACCA	CTTTTGTGAC	TGTCACCCAT	GAGGTATGGA	AGGAGCAGGC	ACTĢGCCTGA	1140
15	GCATGCAGCC	TGGAGAGTGT	TCTTGTCTCT	AGCAGCTGGT	TGGGGACTAT	ATTCTGTCAC	1200
10	TGGAGTTTTG	AATGCAGGGA	CCCCGCATTC	CCATGGTTGT	GCATGGGGAC	ATCTAACTCT	1260
	GGTCTGGGAA	GCCACCCACC	CCAGGGCAAT	GCTGCTGTGA	TGTGCCTTTC	CCTGCAGTCC	1320
20	TTCCATGTGG	GAGCAGAGGT	GTGAAGAGAA	TTTACGTGGT	TGTGATGCCA	AAATCACAGA	1380
	ACAGAATTTC	ATAGCCCAGG	CTGCCGTGTT	GTTTGACTCA	GAAGGCCCTT	CTACTTCAGT	1440
25	TTTGAATCCA	CAAAGAATTA	AAAACTGGTA	ACACCACAGG	CTTTCTGACC	ATCCATTCGT	1500
	TGGGTTTTCC	ATTTGACCCA	ACCCTCTGCC	TACCTGAGGA	GCTTTCTTTG	GAAACCAGGA	1560
	TGGAAACTTC	TTCCCTGCCT	TACCTTCCTT	TCACTCCATT	CATTGTCCTC	TCTGTGTGCA	1620
30	ACCTGAGCTG	GGAAAGGCAT	TTGGATGCCT	CTCTGTTGGG	GCCTGGGGCT	GCAGAACACA	1680
	CCTGCGTTTC	ACTGGCCTTC	ATTAGGTGGC	CCTAGGGAGA	TGGCTTTCTG	CTTTGGATCA	1740
35	CTGTTCCCTA	GCATGGGTCT	TGGGTCTATT	GGCATGTCCA	TGGCCTTCCC	AATCAAGTCT	1800
	CTTCAGGCCC	TCAGTGAAGT	TTGGCTAAAG	GTTGGTGTAA	AAATCAAGAG	AAGCCTGGAA	1860
	GACATCATGG	ATGCCATGGA	TTAGCTGTGC	AACTGACCAG	CTCCAGGTTT	GATCAAACCA	1920
40	AAAGCAACAT	TTGTCATGTG	GTCTGACCAT	GTGGAGATGT	TTCTGGACTT	GCTAGAGCCT	1980
	GCTTAGCTGC	ATGTTTTGTA	GTTACGATTT	TTGGAATCCC	ACTTTGAGTG	CTGAAAGTGT	2040
45	AAGGAAGCTT	TCTTCTTACA	CCTTGGGCTT	GGATATTGCC	CAGAGAAGAA	ATTTGGCTTT	2100
,,,	TTTTTTNCTT	AATGGACAAG	AGACAGTTGC	TGTTCTCATG	TTCCAAGTCT	GAGAGCAACA	2160
	GACCCTCATC	ATCTGTGCCT	GGAAGAGTTC	ACTGTCATTG	AGCAGCACAG	CCTGAGTGCT	2220
50	GGCCTCTGTC	AACCCTTATT	CCACTGCCTT	ATTTGACAAG	GGGTTACATG	CTGCTCACCT	2280
	TACTGCCCTG	GGATTAAATC	AGTTACAGGC	CAGAGTCTCC	TTGGAGGGCC	TGGAACTCTG	2340
55	AGTCCTCCTA	TGAACCTCTG	TAGCCTAAAT	GAAATTCTTA	AAATCACCGA	TGGAACCAAA	2400
55	ааааааааа	ААААААААА	АААААААА	NAAAA			2435

(2) INFORMATION FOR SEQ ID NO: 108:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 805 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 108:

10	(71)	SEQUENCE L	DESCRIPTION.	. SEQ ID NO	. 100.		
10	ATGAAACTTA	AGAATTGAAT	TGGAAAGACT	TCTCAAAGAG	AATTGTATGT	AACGATGTTG	60
	TATTGATTT	TAAGAAAGTA	ATTTAATTTG	TAAAACTTCT	GCTCGTTTAC	ACTGCACATT	120
15	GAATACAGGT	AACTAATTGG	AAGGAGAGGG	GAGGTCACTC	TTTTGATGGT	GGCCCTGAAC	180
	CTCATTCTGG	TTCCCTGCTG	CGCTGCTTGG	TGTGACCCAC	GGAGGATCCA	CTCCCAGGAT	240
20	GACGTGCTCC	GTAGCTCTGC	TGCTGATACT	GGGTCTGCGA	TGCAGCGGCG	TGAGGCCTGG	300
20	GCTGGTTGGA	GAAGGŢCACA	ACCCTTCTCT	GTTGGTCTGC	CTTCTGCTGA	AAGACTCGAG	360
	AACCAACCAG	GGAAGCTGTC	CTGGAGGTCC	CTGGTCGGAG	AGGGACATAG	AATCTGTGAC	420
25	CTCTGACAAC	TGTGAAGCCA	CCCTGGGCTA	CAGAAACCAC	AGTCTTCCCA	GCAATTATTA	480
	CAATTCTTGA	ATTCCTTGGG	GATTTTTTAC	TGCCCTTTCA	AAGCACTTAA	GTGTTAGATC	540
30	TAACGTGTTC	CAGTGTCTGT	CTGAGGTGAC	TTAAAAAATC	AGAACAAAAC	TTCTATTATC	600
30	CAGAGTCATG	GGAGAGTACA	CCCTTTCCAG	GAATAATGTT	TTGGGAAACA	CTGAAATGAA	660
	ATCTTCCCAG	TATTATAAAT	TGTGTATTTA	AAAAAAAGAA	ACTTTTCTGA	ATGCCTACTG	720
35	GCGGTGTATA	CCAGGCAGTG	TGCCAGTTTA	AAAAGATGAA	AAAGAATAAA	AACTITIGAG	780
	GAACAAAAAA	АААААААА	AAATT				805

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(2) INFORMATION FOR SEQ ID NO: 109:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1166 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 109:

	GGCACGAGAG	GCGCCAGTCG	CAGGTGTGCT	GCTGAGGCGT	GAGAATGGCG	TCCCGCGGCC	60
55	GGCGTCCGGA	GCATGGCGGA	CCCCAGAGC	TGTTTTATGA	CGAGACAGAA	GCCCGGAAAT	120
	ACGTTCGCAA	CTCACGGATG	ATTGATATCC	AGACCAGGAT	GGCTGGGCGA	GCATTGGAGC	180
	TTCTTTATCT	GCCAGAGAAT	AAGCCCTGTT	ACCTGCTGGA	TATTGGCTGT	GGCACTGGGC	240
60	TGAGTGGAAG	TTATCTGTCA	GATGAAGGGC	ACTATTGGGT	GGGCCTGGAT	ATCAGCCCTG	300

	CCATGCTGGA	TGAGGCTGTG	GACCGAGAGA	TAGAGGGAGA	CCTGCTGCTG	GGGGATATGG	360
5	GCCAGGGCAT	CCCATTCAAG	CCAGGCACAT	TTGATGGTTG	CATCAGCATT	TCTGCTGTGC	420
J	CCATGCTGGA TGAGGCTGTG GACCGAGAGA TAGAGGGAGA CCTGCTGCTG GGGGATATGG GCCAGGGCAT CCCATTCAAG CCAGGCACAT TTGATGGTTG CATCAGCATT TCTGCTGTGC AGTGGCTCTG TAATGCTAAC AAGAAGTCTG AAAACCCTGC CAAGCGCCTG TACTGCTTTT TTGCTTCTCT TTTTTCTGTT CTCGTCCGGG GATCCCGAGC TGTCCTGCAG CTGTACCCTG AGAACTCAGA GCAGTTGGAG CTGATCACAA CCCAGGCCAC AAAGGCAGGC TTCTCCGGTG GCATGGTGGT AGACTACCCT AACAGTGCCA AAGCAAAGAA ATTCTACCTC TGCTTGTTTT CTGGGCCTTC GACCTTTATA CCAGAGGGGC TGAGTGAAAA TCAGGATGAA GTTGAACCCA GGGAGTCTGT GTTCACCAAT GAGAGGTTCC CATTAAGGAT GTCGAGGCGG GGAATGGTGA GGAAGAGTCG GGCATGGGTG CTGGAGAAGA AGGACCGGCA CAGGCGCCAG GGCAGGGAAG TCAGACCTGA CACCCAGTAC ACCGGCCGCA AGCGCAAGCC CCGCTTCTAA GTCACCACGC GGTTCTGGAA AGGCACTTGC CTCTGCACTT TTCTATATTG TTCAGCTGAC AAAGTAGTAT TTTAGAAAAA TCTTAAAGTT ATAAAAATGT TTTCTGCAGT AAAAAAAAAA	480					
	TTGCTTCTCT	TTTTTCTGTT	CTCGTCCGGG	GATCCCGAGC	TGTCCTGCAG	CTGTACCCTG	540
10	AGAACTCAGA	GCAGTTGGAG	CTGATCACAA	CCCAGGCCAC	AAAGGCAGGC	TTCTCCGGTG	600
	GCATGGTGGT	AGACTACCCT	AACAGTGCCA	AAGCAAAGAA	ATTCTACCTC	TGCTTGTTTT	660
15	CTGGGCCTTC	GACCTTTATA	CCAGAGGGGC	TGAGTGAAAA	TCAGGATGAA	GTTGAACCCA	720
	GGGAGTCTGT	GTTCACCAAT	GAGAGGTTCC	CATTAAGGAT	GTCGAGGCGG	GGAATGGTGA	780
	GGAAGAGTCG	GGCATGGGTG	CTGGAGAAGA	AGGAGCGGCA	CAGGCGCCAG	GGCAGGGAAG	840
20	TCAGACCTGA	CACCCAGTAC	ACCGGCCGCA	AGCGCAAGCC	CCGCTTCTAA	GTCACCACGC	900
	GGTTCTGGAA	AGGCACTTGC	CTCTGCACTT	TTCTATATTG	TTCAGCTGAC	AAAGTAGTAT	960
25	TTTAGAAAAG	TTCTAAAGTT	ATAAAAATGT	TTTCTGCAGT	AAAAAAAAAG	TTCTCTGGGC	1020
23	CGGGCGTGGT	GGCTCACANC	TGTAATCCCA	GCACCTTGGG	AGGCTGAGGT	GGGAGGATCA	1080
	TTTGAGGCCA	GGAGTTTGAG	ACCTGCCTGG	GCAACATAAT	GAAACTTCCT	TTCCAGGGAG	1140
30	AAAAAAA	АААААААА	ACTCGA				1166

35 (2) INFORMATION FOR SEQ ID NO: 110:

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(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 586 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 110:

45 AGAGCGGACG AAGCTGGATA ACAGGGGACC GATGATGTGG CGACCATCAG TTCTGCTGCT 60 TCTGTTGCTA CTGAGGCACG GGGCCCAGGG GAAGCCATCC CCAGACGCAG GCCCTCATGG 120 · CCAGGGGAGG GTGCACCAGG CGGCCCCCT GAGCGACGCT CCCCATGATG ACGCCCACGG 180 50 GAACTTCCAG TACGACCATG AGGCTTTCCT GGGACGGGAA GTGGCCAAGG AATTCGACCA 240 ACTCACCCCA GAGGAAAGCC AGGCCCGTCT GGGGCGGATC GTGGACCGCA TGGACCGCGC .. 300 55 GGGGGACGCC GACGCTGGG TGTCGCTGGC CGAGCTTCGC GCGTGGATCG CGCACACGCA 360 GCAGCGGCAC ATACGGGACT CGGTGAGCGC GGCCTGGGAC ACGTACGACA CGGACCGCGA 420 CGGGCGTGTG GGTTGGGAGG AGCTGCGCAA CGYCACCTAT GGCCACTASG SGCCCGKTGA 480 60

260

	AGAATTTCAT GACGTGGAGG ATGCAGAGAC YTACAAAAAG ATGCTGGYTC GGGACGAGCG	540
	GCGTTTCCGG GTGGCCGACC AGGATGGGGA CTCGATGGCC ACTCGA	586
5		
	(2) INFORMATION FOR SEQ ID NO: 111:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1134 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 111:	
	ACCCATTGAG CAGAAGGAGG CCAGGTGGGA AAGCTCCTGG GAAGAGCAGC CAGACTGGAC	60
20	ACTGGGCTGC TTGAGTCCTG AGTCACAATT CAGAATTCCT GGGCTCCCTG GGTGCATTCT	120
	ATCATTCCAG TTGAAAGTTT GCTTCCTTCC AGTCATGTGG CTCTTCATTC TACTCTCCTT	180
25	GGCTCTCATT TCAGATGCCA TGGTCATGGA TGAAAAGGTC AAGAGAAGTT TGTGCTGGAC	240
23	ACGGCTTCTG CCATCTGCAA CTACAATGCC CAYTACAAGA ATCACCCCAA ATACTGGTGC	300
	CGAGGYTATT TCCGTGAYTA CTGCAACATC ATCGCCTTCT CCCCTAACAG CACCAATCAT	360
30	GTGGCCCTGA AGGACACAGG GAACCAGCTC ATTGTCACTA TGTCCTGCCT GAACAAANAA	420
	GACACGGGCT GGTACTGGTG TGGCATCCAR CGGGACTTTG CMAGGGATGA CATGGATTTT	480
35	ACAGAGCTGA TTGTAACTGA CGACAAAGGA ACCCTGGCCA ATGACTTTTG GTCTGGGAAA	540
33	GACCTATCAG GCAACAAAAC CAGAAGCTGC AAGGCTCCCA AAGTTGTCCG CAAGCTGACC	600
	GCTCCAGGAC GTCCATTCTC ATCATTTGCA TACTGATCAC GGGTTTGGGA ATCATCTCTG	660
40	TAATCAGTCA TTTGACCAAA AGGAGGAGAA GTCAAAGGAA TAGAAGGGTA GGCAACACTT	720
	TGAAGCCCTT CTCGCGTGTC CTGACTCCAA AGGAAATGGC TCCTACTGAA CAGATGTGAC	780
45	TGAAGWITTT TITAATITAG TINCATAAAG TGATGNCTAC AACAGAWTAA TCACCCATGA	840
73	CAACTGGCCC CACACCTCAG AGACTGATTC TGATCTCCCA GGAATTCTGA AGGACCCTCT	900
	ATCCTTGACA ACAATCATTT GCAGCCAGGT AGCAACGGCR GTAGTCAGAG GAGCTATGAT	960
50	AGACCACACC CAAGCAAGGC TGCCCTCAAA TAACATCTCA AGATCTTAGT TCTTATGCAT	1020
	TCCATCAGTC AGAAGTGAAG AAGAGGTGGA GAATCTKGAT TGGGGACCAG GAAATCACTT	1080
55	GTATTITGTT AGCCAATAAA TTCCTAGCCA GTGTTGAATG AAAAAAAAAA	1134

⁽²⁾ INFORMATION FOR SEQ ID NO: 112:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1333 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 112:

10	CACTTTAAAG	CTCTGCTGAG	GGAGTTCGGA	GCCCAGGCTT	TCAGGCGACC	TCTGCCCTCC	60
	CTGCCTCTCC	TCACCCTCCC	TCTCTTCCTG	CAGGGCCTGG	GAAGGGCTTT	GAGGGAGCCT	120
	GGGAGCCATG	TGAAGAGGG	CACGCCTGGG	CTGTCCCACA	GTTTAGATCC	AGTTGGAGGT	180
15	TCTCCCTGGC	TCCTGCAGGC	CTGCGGGGAT	CTCTCCCCAC	TTCAGGCCTC	CGGCAGCTGC	240
	CTGCCCTCTT	GTCTGTGCTT	CAGCCCTGCA	CAAAAGCAGC	TTGGTGACAC	CACTCAGCCA	300
20	CCCAGAGTAC	GTGTTTACAG	GCTTTCCAGA	TCACCTTCCT	GTGGGGTGAA	CGTAATGAGG	360
	CGGGGCTGGT	CCTTGGAATT	TCCCCTGGAA	AATGGTAACA	GACTCCATCC	TTGACCCGGG	420
	GATGAGCATG	AAGGCATTGT	CCCAAAGGCA	GAGGCCACCG	TGGTAGGAAT	TCCACCAAGG	480
25	CCAGAAGGGA	AAAAGGAAGA	ACCCACCGTG	TCTGGCTGTG	CGGGCCCTGG	GGAGGGTCGT	540
	GAGTGCAGCC	CCTCTCTACT	TCYGTGCCTT	TGTAAAACGT	GTAGATAACC	GCAGTGGTTG	600
30	GCTGAGCCAA	GAACTCTCCT	AAATCAGTGG	CTTTCTCCCC	ACCCCTTGCT	GGGGAGTCAT	660
	TTTTAAAAAA	ATCTGTGGGA	TATAAAATTG	GCCTCCTGCT	GCTTCAGCCT	ACCTCTCCCT	720
	CTGCTGACTT	AATGTCGTGA	TTCTGTTTCT	TCAGATATTT	AAGGCTGTTA	GGTTGTGTGA	780
35	GCCTTGAAGT	GTGTGTGTGT	GTCCCAGCGA	CTGTCCACTG	TCCAGGAGAT	GCATGTCTTT	840
	GTATTGGAGA	TATTTCTGTA	ACTCATTCTC	TTGGTGCTCA	CGATTGCCAT	GGCCATAGGG	900
40	CCACAGTGCC	GTATCTGCTG	CAGACATGAT	TGTTTCTTGT	TCTAGAGGTT	TTCTTGTTTT	960
	CGAATCTTGC	CTGATGAATC	CAGCCAGACC	AAGGGCCTA	GATTTGACCT	CTGTCCTGGG	1020
	CTCCTGGGCC	AGGTGCAGGA	ACATCTGAGG	CCACTCTGCT	GGCCACCTCC	AGTGGGTGCT	1080
45	GACCACAGGA	TGGGCTTTGT	TTACACTCAT	TTTCACCCTG	ATTCTTGCCC	CCACTTTCAT	1140
	AAAAGAAACT	TCAAAATGCT	GACGCTTTGG	AGAGTAAGAA	AATCAATCTT	GGCTGGGCAC	1200
50	GGTGGCTCCT	GCCTGTGATC	CTAGCACTTT	GGGAGGCTGA	AGCTGAAGGA	TCACTTGAGC	1260
	TCAGGAGTTG	GAGACCAACC	CTGGCAACAT	AACAAGACCC	TGTCTCTACA	АААААААА	1320
	АААААААСТ	CGA		•.			1333

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⁽²⁾ INFORMATION FOR SEQ ID NO: 113:

⁽i) SEQUENCE CHARACTERISTICS:

WO 98/42738 PCT/US98/05311

262

(A) LENGTH: 1015 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 113:

						•	
	GGCACGAGCG	GCACGAGCGG	CACGAGGTGA	CTTCAAGTGT	CGGATCTTTT	CAGCCTACAT	60
10	CAAGGAGGTG	GAGGAACGGC	CGGCACCCAC	CCCGTGGGCT	CCAAGATGCC	CTTTGGGGAA	120
	CTGATGTTCG	AATCCAGCAG	TAGCTGCGGC	TGGGTACATG	GCGTCTGTTT	CTCAGCCAGC	180
15	GGGAGCCGCG	TGGCCTGGGT	AAGCCACGAC	AGCACCGTCT	GCCTGGCTGA	TGCCGACAAG	240
13	AAGATGGCCG	TCGCGACTCT	GGCCTCTGAA	ACACTACCAC	TGCTGGCGCT	GACCTTCATC	300
	ACAGACAACA	GCCTGGTGGC	AGCGGGCCAC	GACTGCTTCC	CGGTGCTGTT	CACCTATGAC	360
20	GCCGCCGCGG	GGATGCTGAG	CTTCGGCGGG	CGGCTGGACG	TTCCTAAGCA	GAGCTCGCAG	420
	CGTGGCTTGA	CGGCCCGCGA	GCGCTTCCAG	AACCTGGACA	AGAAGGCGAG	CTCCGAGGGT	480
25	GGCACGGCTG	ceeecece	CCTAGACTCG	CTGCACAAGA	ACAGCGTCAG	CCAGATCTCG	540
23	GTGCTCAGCG	GCGGCAAGGC	CAAGTGCTCG	CAGTTCTGCA	CCACTGGCAT	GGATGGCGGC	600
	ATGAGTATCT	GGGATGTGAA	GAGCTTGGAG	TCAGCCTTGA	AGGACCTCAA	GATCAAATGA	660
30	CCTGTGAGGA	ATATGTTGCC	TTCATCCTAG	CTGCTGGGGA	AGCGGGGAGA	GGGTCAGGG	720
	AGGCTAATGG	TTGCTTTGCT	GAATGTTTCT	GGGTACCAA	TACGAGTTCC	CATAGGGGCT	780
35	GCTCCCTCAA	AAAGGGAGGG	GACAGATGGG	GAGCTTTTCT	TACCTATTCA	AGGAATACGT	840
33	GCCTTTTTCT	TAAATGCTTT	CATTTATTGA	ааааааааа	AAATGCCCCC	AAAGCACTAT	900
	GCTGGTCATG	AACTGCTTCA	AAATGTGGAG	GTAATAAAAT	GCAACTGTGT	АААААААА	960
40	АААААААА	АААААААА	ааааааааа	ааааааааа	ААААААААА	AACNC	1015

45 (2) INFORMATION FOR SEQ ID NO: 114:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1076 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 114:

55	GGCACGAGGG	GAAAGCCATG	CTCCCAGGAC	TCCTTCCTTG	CAGCCTTAAA	TCGGTCTGTA	60
	CGGAAAATTC	CGCGCCTTAG	AAACCCACGC	TTGGGTGTAA	CTTATTATTG	TTCTTCCTGA	120
60	CCTACTTCCT	GTTTATCACT	TCCGGGTTCA	TCATTTTGGC	ATTTCGGTGA	TCGGGTTGGA	180

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	ACTATTGAAG	CCCGCTTTCA	GGTTCTTTTC	CCCATTTTCC	CTTTGAAAGG	AAGACTTCTG	240
	GCTTCTCCTA	AATCTCCGTT	CTCTGGGTAA	GGGGAGTCCA	AGCCTCTGTC	ATGAGGAACG	300
5	GAAATGCGAG	GCCTCGGGT	GTTACTCTAA	AATCCGCCCT	CAGCTTGCAC	GCCGGAAGCT	360
	GCGATTCCTG	CAGCGGAAGA	GGCGTGATCT	GGCCTTCGAC	TCGCTATGTC	CACTAACAAT	420
10	ATGTCGGACC	CACGGAGGCC	GAACAAAGTG	CTGAGGTACA	AGCCCCCGCC	GAGCGAATGT	480
10	AACCCGGCCT	TGGACGACCC	GACGCCGGAC	TACATGAACC	TGCTGGGCAT	GATCTTCAGC	540
	ATGTGCGGCC	TCATGCTTAA	GCTGAAGTGG	TGTGCTTGGG	TCGCTGTCTA	CTGCTCCTTC	600
15	ATCAGCTTTG	CCAACTCTCG	GAGCTCGGAG	GACACGAAGC	AAATGATGAG	TAGCTTCATG	660
	CTGTCCATCT	CTGCCGTGGT	GATGTCCTAT	CTGCAGAATC	CTCAGCCCAT	GACGCCCCCA	720
20	TGGTGATACC	AGCCTAGAAG	GGTCACATTT	TGGACCCTGT	СТАТССАСТА	GCCTGGGCT	780
20	TTGGCTGCTA	AACCTGCTGC	CTTCAGCTGC	CATCCTGGAC	TTCCCTGAAT	GAGGCCGTCT	840
	CGGTGCCCCC	AGCTGGATAG	AGGGAACCTG	GCCCTTTCCT	AGGGAACACC	CTAGGCTTAC	900
25	CCCTCCTGCC	TCCCTTCCCC	TGCCTGCTGC	TGGGGGAGAT	GCTGTCCATG	TTTCTAGGGG	960
	TATTCATTTG	CTTTCTCGTT	GAAACCTGTT	GTTAATAAAG	TTTTTCACTC	TGAAAAAAA	1020
20	, AAAAAAAAA	RAAAACNCGN	GGGGGGCCC	GGAACCCAAT	TCSCCGGATA	GTGAGT	1076
30							

(2) INFORMATION FOR SEQ ID NO: 115:

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(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1487 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 115:

45	CCGCTGCTGA	TAACTATGGC	ATCCCCCGGG	CCTGCAGGAA	TTCGGCACGG	AGCTACGGCG	60
43	CCGCCTGGCT	ACCAC CAGGCTCATC GTGAACCTGT CCCAGACCTA CATGGCCATG TACCTCACCT 12 13 14 15 16 16 17 17 18 18 18 18 18 18 18 18	120				
	GCATCAGCGA	CCAGCTGGGG	GGCCAGGACG	TGCCCGTGTT	CCGGAACCTG	TCCCTGCTGG	180
50	TGGTGGGTGT	CGGCGCCGTG	TTCTCACTGC	TATTCCACCT	GGGCACCCGG	GAGAGGCGCC	240
	GGCCGCATGC	GGASGAGCCA	GGCGAGCACA	CCCCCTGTT	GCCCCTGCC	ACGGCCCAGC	300
55	CCCTGCTGCT	CTGGAAGCAC	TGGCTCCGGG	AGCSGGCTTT	CTACCAGGTG	GGCATACTGT	360
	ACATGACCAC	CAGGCTCATC	GTGAACCTGT	CCCAGACCTA	CATGGCCATG	TACCTCACCT	420
	ACTCGCTCCA	CCTGCCCAAG	AAGTTCATCG	CGACCATTCC	CCTGGTGATG	TACCTCAGCG	480
60	GCTTCTTGTC	CTCCTTCCTC	ATGAAGCCCA	TCAACAAGTG	CATTGGGAGG	AACATGACCT	540

	ACTTCTCAGG	CCTCCTGGTG	ATCCTGGCCT	TTGCCGCCTG	GGTGGCGCTG	GCGGAGGGAC	600
5	TGGGTGTGGC	CGTGTACGCA	GCGGCTGTGC	TGCTGGGTGC	TGGCTGTGCC	ACCATCCTCG	660
J	TCACCTCGCT	GGCCATGACG	GCCGACCTCA	TCGGTCCCCA	CACGAACAGC	GGAGCKTTCG	720
	TGTACGGCTC	CATGAGCTTC	TTGGATAAGG	TGGCCAATGG	GCTGGCAGTC	ATGGCCATCC	780
0	AGAGCCTGCA	CCCTTGCCCC	TCAGAGCTCT	GCTGCAGGGC	CTGCGTGAGC	TTTTACCACT	840
	GGGCGATGGT	GGCTGTGACG	GCCGCCTCG	GCGTGGCCGC	TGCCCTGTGT	CTCTGTAGCC	900
15	TCCTGCTGTG	GCCGACCCGC	CTGCGACGCT	GATGAGACCT	GCACGCANTG	GCTCACAGCA	960
	GCACGATTTG	TGACAGCCCG	AGGCGGAGAA	CACCGAACAC	CCAGTGAAGG	TGAGGGGATC	1020
	AGCACGGCGC	GGCCACCCAC	GCACCCACGC	GCTGGAATGA	GACTCAGCCA	CAAGGAGGTG	1080
20	CGAAGCTCTG	ACCCAGGCCA	CAGTGCGGAT	GCACCTTGAG	GATGTCACGC	TCAGTGAGAG	1140
	ACACCAGACA	CGC GGCCACCCAC GCACCCACGC GCTGGAATGA GACTCAGCCA CAAGGAGGTG CTG ACCCAGGCCA CAGTGCGGAT GCACCTTGAG GATGTCACGC TCAGTGAGAG GACA CAGAAGGGTA CGCTGTGATC CCACTTCTAT GAAATGTCCA GGACAGACCA	1200				
25	ATCCACAGAA	TCAGGGAGAG	GATTCGTGGG	TGCCGGGACT	GGGGAGGGG	ACCTGGGGGT	1260
••	GACTAGGTGA	CATAATGGGG	ACAGGGCTGC	CTTCTGGGTG	ATGAGAATGT	TCTGGAATCA	1320
	GATGGGATGG	CTGCACGGCG	TGGTGAAGGT	ACTGAACGCC	ACCTCACTGT	AAGACGGTAG	1380
30	ATTTTGTATT	TTACCACAAT	AAACAAAACA	AAACAAAACC	АААААААА	AAAAAAAA	1440
	AAAAAAAAGG	AATTCGATAT	CAAGCTTATC	GATACCGTCG	ACCTCGA		1487

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(2) INFORMATION FOR SEQ ID NO: 116:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1350 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 116:

GGCACGAGTG CGCANGCGTG GGGCTCTCTC CTTGTCAGTC GGCGCCGCGT GCGGGCTGGT 60 GGCTCTGTGG CAGCGGCGGC GGCAGGACTC CGGCACTATG AGCGGCTTCA GCACCGAGGA 120 50 GCGCGCCGCG CCTTCTCCCT GGAGTACCGA GTCTTCCTCA AAAATGAGAA AGGACAATAT 180 ATATCTCCAT TTCATGATAT TCCAATTTAT GCAGATAAGG ATGTGTTTCA CATGGTAGTT 240 55 GAAGTACCAC GCTGGTCTAA TGCAAAAATG GAGATTGCTA CAAAGGACCC TTTAAACCCT 300 ATTAAACAAG ATGTGAAAAA AGGAAAACTT CGCTATGTTG CGAATTTGTT CCCGTATAAA 360 GGATATATCT GGAACTATGG TGCCATCCCT CAGACTTGGG AAGACCCAGG GCACAATGAT 420 60

480

	AAACATACTG GCTGTTGTGG TGACAATGAC CCAATTGATG TGTGTGAAAT TGGAAGCAAG	480
	GTATGTGCAA GAGGTGAAAT AATTGGCGTG AAAGTTCTAG GCATATTGGC TATGATTGAC	540
5	GAAGGGGAAA CCGACTGGAA AGTCATTGCC ATTAATGTGG ATGATCCTGA TGCAGCCAAT	600
	TATAATGATA TCAATGATGT CAAACGGCTG AAACCTGGCT ACTTAGAAGC TACTGTGGAC	660
10	TGGTTTAGAA GGTATAAGGT TCCTGATGGA AAACCAGAAA ATGAGTTTGC GTTTAATGCA	720
10.	GAATTTAAAG ATAAGGACTT TGCCATTGAT ATTATTAAAA GCACTCATGA CCATTGGAAA	780
	GCATTAGTGA CTAAGAAAAC GAATGGAAAA GGAATCAGTT GCATGAATAC AACTTTGTCT	840
15	GAGAGCCCCT TCAAGTGTGA TCCTGATGCT GCCAGAGCCA TTGTGGATGC TTTACCACCA	900
	CCCTGTGAAT CTGCCTGCAC AGTACCAACA GACGTGGATA AGTGGTTCCA TCACCAGAAA	960
20	AACTAATGAG ATTTCTCTGG AATACAAGCT GATATTGCTA CATCGTGTTC ATCTGGATGT	1020
5 G 10 G 15 G 20 A 20 A 35 G 40 G 55 G 55 G 66 G 55 G 67 G 68	ATTAGAAGTA AAAGTAGTAG CTTTTCAAAG CTTTAAATTT GTAGAACTCA TCTAACTAAA	1080
	GTAAATTCTG CTGTGACTAA TCCAATATAC TCAGAATGTT ATCCATCTAA AGCATTTTTC	1140
25	ATATCTCAAC TAAGATAACT TTTAGCACAT GCTTAAATAT CAAAGCAGTT GTCATTTGGA	1200
	AGTCACTTGT GAATAGATGT GCAAGGGGAG CACATATTGG ATGTATATGT TACCATATGT	1260
30	TAGGAAATAA AATTATTTTG CTGAAAAAAA AAAAAAAAAA	1320
50	CCCCATTIGG CCCTTIGGGG GGNGGTTTTA	1350
35	(2) INFORMATION FOR SEQ ID NO: 117:	
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2527 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 117:	
43	CTCTTGCTAC CTTCCCGGCG CAGAGAACCC CGGCTGCTCA GCGCGCTCCG GGGTCATGGA	60
	GATCCCCGGG AGCCTGTGCA AGAAAGTCAA GCTGAGCAAT AACGCGCAGA ACTGGGGAAT	120
50	GCAGAGAGCA ACCAATGTCA CCTACCAAGC CCATCATGTC AGCAGGAACA AGAGAGGTCA	180
	GGTGGTGGGG ACCAGAGGTG GCTTTCGTGG TTGCACAGTT TGGCTAACAG GCTTGTCTGG	240
55	AGCGGGAAAG ACTACTGTGA GCATGGCCTT GGAGGAGTAC CTGGTTTGTC ATGGTATTCC	300
J J	ATGCTACACT CTGGATGGTG ACAATATTCG TCAAGGTCTC AATAAAAATC TTGGCTTTAG	360
	TCCTGAAGAC AGAGAAGAGA ATGTTCGACG CATCGCAGAA GTTGCTAAAC TGTTTGCAGA	420

TGCTGGCTTA GTGTGCATCA CAAGTTTCAT ATCACCTTAC ACTCAGGATC GCAACAATGC

	AAGGCAAATT	CATGAAGGTG	CAAGTTTACC	GTTTTTTGAA	GTATTTGTTG	ATGCTCCTCT	540
5	GCATGTTTGT	GAACAGAGGG	ATGTCAAAGG	ACTCTACAAA	AAAGCCCGGG	CAGGAGAAAT	600
J	TAAAGGTTTC	ACTGGGATCG	ATTCTGAATA	TGAAAAGCCA	GAGGCCCCTG	AGTTGGTGCT	660
	GAAAACAGAC	TCCTGTGATG	TAAATGACTG	TGTCCAGCAA	GTTGTGGAAC	TTCTACAGGA	720
10	ACGGGATATT	GTACCTGTGG	ATGCATCTTA	TGAAGTAAAA	GAACTATATG	TGCCAGAAAA	780
	TAAACTTCAT	TTGGCAAAAA	CAGATGCGGA	AACATTACCA	GCACTGAAAA	TTAATAAAGT	840
15	GGATATGCAG	TGGGTGCAGG	TTTTGGCAGA	AGGTTGGGCA	ACCCCATTGA	ATGGCTTTAT	900
13	GAGAGAGAGG	GAGTACTTGC	AGTGCCTTCA	TTTTGATTGT	CTTCTGGATG	GAGGTGTCAT	960
	TAACTTGTCA	GTACCTATAG	TTCTGACTGC	GACTCATGAA	GATAAAGAGA	GGCTGGACGG	1020
20	CTGTACAGCA	TTTGCTCTGA	TGTATGAGGG	CCGCCGTGTG	GCCATTCTTC	GCAATCCAGA	1080
	GTTTTTTGAG	CACAGGAAAG	AGGAGCGCTG	TGCCAGACAG	TGGGGAACGA	CATGCAAGAA	1140
25	CCACCCCTAT	ATTAAGATGG	TGATGGAACA	AGGAGATTGG	CTGATTGGAG	GAGATCTTCA	1200
	AGTCTTGGAT	CGAGTTTATT	GGAATGATGG	TCTTGATCAG	TATCGTCTTA	CTCCTACTGA	1260
	GCTAAAGCAG	AAATTTAAAG	ATATGAATGC	TGATGCTGTC	TTTGCATTTC	AACTACGCAA	1320
30	CCCAGTGCAC	AATGGACATG	CCCTGTTAAT	GCAGGATACC	CATAAGCAAC	TTCTAGAGAG	1380
	GGGCTACCGG	CGCCCTGTCC	TCCTCCTCCA	CCCTCTGGGT	GGCTGGACAA	AGGATGACGA	1440
35	TGTTCCTTTG	ATGTGGCGTA	TGAAGCAGCA	TGCTGCAGTG	TTGGAGGAAG	GAGTTCTGAA	1500
	TCCTGAGACG	ACAGTGGTGG	CCATCTTCCC	ATCTCCCATG	ATGTATGCTG	GACCAACTGA	1560
	GGTCCAGTGG	CATTGCAGAG	CACGGATGGT	TGCAGGAGCC	AACTTTTACA	TTGTTGGACG	1620
40	AGACCCTGCT	GGCATGCCTC	ATCCAGAAAC	AGGGAAGGAT	CTTTATGAGC	CAAGTCATGG	1680
	TGCCAAAGTG	CTGACGATGG	CCCCTGGTTT	AATCACTTTG	GAAATAGTTC	CCTTTCGAGT	1740
45	TGCAGCTTAC	AACAAGAAAA	AGAAGCGTAT	GGACTACTAT	GACTCTGAAC	ACCATGAAGA	1800
.5	CTTTGAATTT	ATTTCAGGAA	CACGAATGCG	CAAACTTGCT	CGAGAAGGCC	AGAAACCACC	1860
	TGAAGGTTTC	ATGGCTCCCA	AGGCTTGGAC	CGTGCTGACA	GAATACTACA	AATCCTTGGA	1920
50	GAAAGCTTAG	GCTGTTAACC	CAGTCACTCC	ACCTTTGACA	CATTACTAGT	AACAAGAGGG	1980
	GACCACATAG	TCTCTGTTGG	CATTTCTTTG	TGGTGTCTGT	CTGGACATGC	TTCCTAAAAA	2040
55	CAGACCATTT	TCCTTAACTT	GCATCAGTTT	TGGTCTGCCT	TATGAGTTCT	GTTTTGAACA	2100
JJ	AGTGTAACAC	ACTGATGGTT	TTAATGTATC	TITTCCACTT	ATTATAGTTA	TATTCCTACA	2160
	ATACAATTTT	AAAATTGTCT	TTTTATATTA	TATTTATGCT	TCTGTGTCAT	GATTTTTCA	2220
60	AGCTGTTATA	TTAGTTGTAA	CCAGTAGTAT	' TCACATTAAA	الململيك كالملائكية	י יייייר רר רייייים	2280

1020

1080

	AAAAAAGAAA AAAATTACCA AACAATAAAC TTGGCTAGAC CTTGTTTTGA GGATTTTACA	2340
5	AGACCTTTGT AGCGATTAGA TTTTTTTTCT ACATTGAAAA TAGAAACTGC TTCCTTTCTT	2400
J	CTITCCAGTC AGCTATTGGT CTITCCAGCT GTTATAATCT AAAGTATTCT TATGATCTGT	2460
	GTAAGCTCTG AATGAACTTC TTTACTCAAT AAAATTAATT TTTTGGCTTC TTAAAAAAAA	2520
10	AAAAAA	2527
	•	
15	(2) INFORMATION FOR SEQ ID NO: 118:	
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1098 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 118:	
25	CGCATCACAG ACAACCCAGA AGGAAAATGG TTGGGCAGAA CAGCAAGGGG TTCATATGGC	60
	TATATTAAAA CAACTGCTGT AGAGATTNNC TATGATTCTT TGAAACTGAA AAAAGACTCT	120
30	CTTGGTGCCC CTTCAAGACC TATTGAAGAT GACCAAGAAG TATATGATGA TGTTGCAGAG	180
50	CAGGATGATA TTAGCAGCCA CAGTCAGAGT GGAAGTGGAG GGATATTCCC TCCACCACCA	240
	GATGATGACA TTTATGATGG GATTGAAGAG GAAGATGCTG ATGATGGTTT CCCTGCTCCT	300
35	CCTAAACAAT TGGACATGGG AGATGAAGTT TACGATGATG TGGATACCTC TGATTTCCCT	360
	GTTTCATCAG CAGAGATGAG TCAAGGAACT AATGTTGGAA AAGCTAAGAC AGAAGAAAAG	420
40	GACCTTAAGA AGCTAAAAAA GCAGRAAAAA GAARAAAAAG ACTTCAGGAA AAAATTTAAA	480
40	TATGATGGTG AAATTAGAGT CCTATATTCA ACTAAAGTTA CAACTTCCAT AACTTCTAAA	540
	AAGTGGGGAA CCAGAGATCT ACAGGTAAAA CCTGGTGAAT CTCTAGAAGT TATACAAACC	600
45	ACAGATGACA CAAAAGTTCT CTGCAGAAAT GAAGAAGGGA AATATGGTTA TGTCCTTCGG	660
	AGTTACCTAG CGGACAATGA TGGAGAGATC TATGATGATA TTGCTGATGG CTGCATCTAT	720
50	GACAATGACT AGCACTCAAC TTTGGTCATT CTGCTGTGTT CATTAGGTGC CAATGTGAAG	780
50	TCTGGATTTT AATTGGCATG TTATTGGGTA TCMAGAAAAT TAATGCACAR AACCACTTAT	840
	TATCATTTGT TATGAAATCC CAATTATCTT TACAAAGTGT TTAAAGTTTG AACATAGAAA	900
55	ATAATCTCTC TGCTTAATTG TTATCTCAGA AGACTACATT AGTGAGATGT AAGAATTATT	960

AAATATTCCA TTICCGCTTT GGCTACAATT ATGAAGAAGT TGAAGGTACT TCTTTTAGAC

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GGGGCCCGG TACCCAAT 1098

5

10

(2) INFORMATION FOR SEQ ID NO: 119:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1679 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 119:

15 TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT 60 CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG 120 20 180 CAGCCGGGAG CCCGCAGCCC GCGCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA 240 CGAGCCCAAG AGCGGCGAGG AGCGCCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG 300 25 CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG 360 CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA 420 30 TTTTGCACTT CAACCAGATG ACGTGTACTA CTGTGGAATA AAGTACATCA AAGATGATGT 480 540 CATCTTAAAT GAGCCCTCTG CAGATGCCCC AGCTGCTCTC TACCAGACAA TTGAAGAAAA 600 TATTAAAATC TTTGAAGAAG AAGAAGTTGA ATTTATCAGT GTGCCTGTCC CAGAGTTTGC 35 AGATAGTGAT CCTGCCAACA TTGTTCATGA CTTTAACAAG AAACTTACAG CCTATTTAGA 660 720 TCTTAACCTG GATAAGTGCT ATGTGATCCC TCTGAACACT TCCATTGTTA TGCCACCCAG 40 AAACCTACTG GAGTTACTTA TTAACATCAA GGCTGGAACC TATTTGCCTC AGTCCTATCT GATTCATGAG CACATGGTTA TTACTGATCG CATTGAAAAC ATTGATCACC TGGGTTTCTT TATTTATCGA CTGTGTCATG ACAAGGAAAC TTACAAACTG CAACGCAGAG AAACTATTAA 900 45 AGGTATTCAG AAACGTGAAG CCAGCAATTG TTTCGCAATT CGGCATTTTG AAAACAAATT 960 1020 TGCCGTGGAA ACTTTAATTT GTTCTTGAAC AGTCAAGAAA AACATTATTG AGGAAAATTA 50 ATATCACAGC ATAACCCCAC CCTTTACATT TTGTGCAGTG ATTATTTTT AAAGTCTTCT 1080 TTCATGTAAG TAGCAAACAG GGCTTTACTA TCTTTTCATC TCATTAATTC AATTAAAACC 1140 ATTACCTTAA AATTTTTTC TTTCGAAGTG TGGTGTCTTT TATATTTGAA TTAGTAACTG 1200 55 . TATGAAGTCA TAGATAATAG TACATGTCAC CTTAGGTAGT AGGAAGAATT ACAATTTCTT 1260 TAAATCATTT ATCTGGATTT TTATGTTTTA TTAGCATTTT CAAGAAGACG GATTATCTAG 1320 60 AGAATAATCA TATATATGCA TACGTAAAAA TGGACCACAG TGACTTATTT GTAGTTGTTA 1380 WO 98/42738 PCT/US98/05311

269

	GTTGCCCTGC	TACCTAGTTT	GTTAGTGCAT	TTGAGCACAC	TTTAATTTTA	TCCTCTAATT	1440
5	AAAATGTGCA	GTATTTTCAG	TGTCAAATAT	ATTTAACTAT	TTAGAGAATG	ATTTCCACCT	1500
5	TTATGTTTTA	ATATCCTAGG	CATCTGCTGT	AATAATATTT	TAGAAAATGT	TTGGAATTTA	1560
	AGAAATAACT	TGTGTTACTA	ATTTGTATAA	CCCATATCTG	TGCAATGGAA	TATAAATATC	1620
10	ACAAAGTTGT	TTAAMWAAAA	АААААААА	аааааааааа	AAAAAAAA	AAAAAAAN	1679

15 (2) INFORMATION FOR SEQ ID NO: 120:

20

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1308 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 120:

25 TTGGCANCNG GGAGAGGGAA AGAGGAGGAA ATGGGGTTTG AGGACCATGG CTTACCTTTC 60 CTGCCTTTGA CCCATCACAC CCCATTTCCT CCTCTTTCCC TCTCCCCGCT GCCAAAAAAA 120 180 AAAAAAAGG AAACGTTTAT CATGAATCAA CAGGGTTTCA GTCCTTATCA AAGAGAGATG 30 240 TGGAAAGAGC TAAAGAAACC ACCCTTTGTT CCCAACTCCA CTTTACCCAT ATTTTATGCA ACACAAACAC TGTCCTTTTG GGTCCCTTTC TTACAGATGG ACCTCTTGAG AAGAATTATC 300 360 35 GTATTCCACG TTTTTAGCCC TCAGGTTACC AAGATAAATA TATGTATATA TAACCTTTAT TATTGCTATA TCTTTGTGGA TAATACATTC AGGTGGTGCT GGGTGATTTA TTATAATCTG 420 480 40 AAAAGCCAGG TATAATGTAA CTTCACCCCA GCCTTTGTAC TAAGCTCTTG ATAGTGGATA 540 TACTCTTTTA AGTTTAGCCC CAATATAGGG TAATGGAAAT TTCCTGCCCT CTGGGTTCCC 600 45 660 CATTITTACT ATTAAGAAGA CCAGTGATAA TTTAATAATG CCACCAACTC TGGCTTAGTT 720 AAGTGAGAGT GTGAACTGTG TGGCAAGAGA GCCTCACACC TCACTAGGTG CAGAGAGCCC 780 AGGCCTTATG TTAAAATCAT GCACTTGAAA AGCAAACCTT AATCTGCAAA GACAGCAGCA 50 AGCATTATAC GGTCATCTTG AATGATCCCT TTGAAATTTT TTTTTTGTTT GTTTGTTTAA 840 ATCAAGCCTG AGGCTGGTGA ACAGTAGCTA CACACCCATA TTGTGTGTTC TGTGAATGCT ... 900 55 AGCTCTCTTG AATTTGGATA TTGGTTATTT TTTATAGAGT GTAAACCAAG TTTTATATTC 960 1020 TGCAATGCGA ACAGGTACCT ATCTGTTTCT AAATAAAACT GTTTACATTC ATTATGGGGT 1080 ATGTATGACC TTCATTTTCC AAGAAATAGA ACTCTAGCTT AGAATTATGG ATGCTCTAAA 60

1200

	ATGTCAGAAT	GGGAACTCTC	CTCGAAGTTC	TCCCAAACTC	AGAGACAGCA	CTGCCTTCTC	1140
	CTAAATGATT	ATTCTTTTCT	CCCTGTTTTC	TGGTATTTTC	TAGGCATCCT	TCTCACCACA	1200
5	GCCATAACCC	TTTTTTACTT	CCATTAGGCC	GTATAACTGG	NGGGACNGCT	GGTCGGTATA	1260
	TAATACTGGT	WCCAACAMAG	GGGTTCTGGA	TGTACACMAG	GTTATCTT		1308
10							
10	(2) INFORMA	MION FOR G	70 TD V0 1	2.1			
		ATION FOR SE				•	
15	(i)	(B) TYP (C) STR	HARACTERIST GTH: 1411 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
20	(xi)	SEQUENCE	DESCRIPTION	: SEQ ID NO	: 121:		
	GGCACAGGAG	CGACCCGGGA	GAAGGAGGC	CAMGAKGCGG	AAGCGGAGGA	GTCTCCAGGA	60
25	GACCCGGGGA	CAGCATCGCC	CAGGCCCCTG	TTTGCAGGCC	TTTCAGATAT	ATCCATCTCA	120
	CAAGACATCC	CCGTAGAAGG	AGAAATCACC	ATTCCTATGA	GATCTCGCAT	CCGGGAGTTT	180
	GACAGCTCCA	CATTAAATGA	ATCTGTTCGC	AATACCATCA	TGCGTGATCT	AAAAGCTGTT	240
30	GGGAAAAAAT	TCATGCATGT	TTTGTACCCA	AGGAAAAGTA	ATACTCTTTT	GAGAGATTGG	300
	GATTTGTGGG	GCCCTTTGAT	CCTTTGTGTG	ACACTCGCAT	TAATGCTGCA	AAGAGACTCT	360
35	GCAGATAGTG	AAAAAGATGG	AGGGCCCCAA	TTTGCAGAGG	TGTTTGTCAT	TGTCTGGTTT	420
	GGTGCAGTTA	CCATCACCCT	CAACTCAAAA	CTTCTTGGAG	GGAACATATC	TTTTTTCAG	480
	AGCCTCTGTG	TGCTGGGTTA	CTGTATACTT	CCCTTGACAG	TAGCAATGCT	GATTTGCCGG	540
40	CTGGTACTTT	TGGCTGATCC	AGGACCTGTA	AACTTCATGG	TTCGGCTTTT	TGTGGTGATT	600
	GTGATGTTTG	CCTGGTCTAT	AGTTGCCTCC	ACAGCTTTCC	TTGCTGATAG	CCAGCCTCCA	660
45	AACCGCAGAG	CCCTAGCTGT	TTATCCTGTT	TTCCTGTTTT	ACTTTGTCAT	CAGTTGGATG	720
	ATTCTCACCT	TTACTCCTCA	GTAAATCAGG	AATGGGAAAT	TAAAAACCAG	TGAATTGAAA	780
	GCACATCTGA	AAGATGCAAT	TCACCATGGA	GCTTTGTCTC	TGGCCCTTAT	TTGTCTAATT	840
50	TTGGAGGTAT	TTGATAACTG	AGTAGGTGAG	GAGATTAAAA	GGGAGCCATA	TAGCACTGTC	900
	ACCCCTTATT	TGAGGAACTG	ATGTTTGAAA	GGCTGTTCTT	TTCTCTCTTA	ATGTCATTTC	960
55	TTTAAAAATA	CATGTGCATA	CTACACACAG	TATATAATGC	CTCCTTAAGG	CATGATGGAG	1020
	TCACCGTGGT	CCATTTGGGT	GACAACCAGT	GACTTGGGAA	GCACATAGAT	ACATCTTACA	1080
	AGTTGAATAG	AGTTGATAAC	TATTTTCAGT	TTTGAGAATA	CCAGTTCAGG	TGCAGCTCTT	. 1140

AAACACATTG CCTTATGACT ATTAGAATAT GCCTCTCTTT TCATAAATAA AAATACATGG

	TCTATATCCA TTTTCTTTTA TTTCTCTCTC TTAAGCTTAA AAAGGCAATG AGAGAGGTTA	1260
5	GGAGTGGGTT CATACACGGA GAATGAGAAA ACATGCATTA ACCAATATTC AGATTTTGAT	1320
5	CAGGGGAAAT TCTAYACTTG TTGCAAAAAA AAAAAAAAA AAACTCGAGG GGGGCCCGGT	1380
	ACCCAATCGC NGTATATGAT CGNAAACAAT C	1411
10		
. ~	(2) INFORMATION FOR SEQ ID NO: 122:	
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 2256 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 122:	
	GCTTTGGCTT TTTTTGGCGG ACTGGGGCGC CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG	60
25	TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA	120
	GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC	180
30	CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC	240
30	TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG	300
	CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG	360
35	AGCGCANGCC GGCCTGGCCT TCAGCTTGTA CCAGGCCATG GCCAAGGACC AGGCAGTGGA	420
	GAACATCCTG GTGTCACCCG TGGTGGTGGC CTCGTCGCTG GGGCTCGTGT CGCTGGGCGG	480
40	CAAGGCGACC ACGGCGTCGC AGGCCAAGGC AGTGCTGAGC GCCGAGCAGC TGCGCGACGA	540
40	GGAGGTGCAC GCCGGCCTGG GCGAGCTGCT GCGCTCACTC AGCAACTCGA CGGCGCGCAA	600
	CGTGACCTGG AAGCTGGGCA GCCGACTGTA CGGACCCAGC TCAGTGAGCT TCGCTGATGA	660
45	CTTCGTGCGC ACAGCAAGCA GCACTACAAC TGCGAGCACT CCAAGATCAA CTTCCGCGAC	720
	AAGCGCAGNG CGCTGCAGTC CATCAACGAG TGGGCCGCGC AGACCACCGA CGGCAAGCTG	780
50	CCCGAGGTCA CCAAGGACGT GGAGCGCACG GACGGCGCCC TGCTAGTCAA CGCCATGTTC	840
30	TTCAAGCCAC ACTGGGATGA GAAATTCCAC CACAAGATGG TCGACAACCG TCGCTTCATG	900
	GTGACTCGGT CCTATACYGT GGGTGTCATG ATGATGCACC GGACAGGCCT CTACAACTAC	960
55	TACGACGACG AGAAGGAAAA GCTGCAAATC GTGGAGATGC CCCTGGCCCA CAAGCTCTCC	1020
	AGCCTCATCA TCCTCATGCC CCATCACGTG GAGCCTCTCG AGCGCCTTGA AAAGCTGCTA	1080
CO	ACCAAAGAGC AGCTGAAGAT CTGGATGGGG AAGATGCAGA AGAAGGCTGT TGCCATCTCC	1140

	TTGCCCAAGG	GTGTGGTGGA	GGTGACCCAT	GACCTGCAGA	AACACCTGGC	TGGGCTGGGC	1200
	CTGACTGAGG	CCATTGACAA	GAACAAGGCC	GACTTRTCAC	GCATGTCAGG	CAAGAAGGAC	1260
5	CTGTACCTGG	CCAGCGTGTT	CCACGCCACC	GCCTTTGAGT	TGGACACAGA	TGGCAACCCC	1320
	TTTGACCAGG	ACATCTACGG	GCGCGAGGAG	CTGCGCANCC	CAAGCTGTTC	TACGCCGACC	1380
10	ACCCCTTCAT	CTTCCTAGTG	CGGGACACCC	AAAGCGGCTC	CCTGCTATTC	ATTGGGCGCC	1440
10	TGGTCCGGCC	TAAGGGTGAC	AAGATGCGAG	ACGAGTTATA	GGGCCTCAGG	GTGCACACAG	1500
	GATGGCAGGA	GGCATCCAAA	GGCTCCTGAG	ACACATGGGT	GCTATTGGGG	TTGGGGGGGA	1560
15	GGTGAGGTAC	CAGCCTTGGA	TACTCCATGG	GGTGGGGGTG	GAAAARCAGA	CCGGGGTTCC	1620
	CGTGTGCCTG	AGCGGACCTT	CCCAGCTAGA	ATTCACTCCA	CTTGGACATG	GGCCCCAGAT	1680
20	ACCATGATGC	TGAGCCCGGA	AACTCCACAT	CCTGTGGGÄC	CTGGGCCATA	GTCATTCTGC	1740
20	CTGCCCTGAA	AGTCCCAGAT	CAAGCCTGCC	TCAATCAGTA	ТТСАТАТТТА	TAGCCAGGTA	1800
	CCTTCTCACC	TGTGAGACCA	AATTGAGCTA	GGGGGTCAG	CCAGCCCTCT	TCTGACACTA	1860
25	AAACACCTCA	GCTGCCTCCC	CAGCTCTATC	CCAACCTCTC	ССААСТАТАА	AACTAGGTGC	1920
	TGCAGCCCCT	GGGACCAGGC	ACCCCCAGAA	TGACCTGGCC	GCAGTGAGGC	GGATTGAGAA	1980
30	GGAGCTCCCA	GGAGGGGCTT	CTGGGCAGAC	TCTGGTCAAG	AAGCATCGTG	TCTGGCGTTG	2040
50	TGGGGATGAA	CTTTTTGTTT	TGTTTCTTCC	TTTTTTAGTT	CTTCAAAGAT	AGGGAGGGAA	2100
	GGGGGAACAT	GAGCCTTTGT	TGCTATCAAT	CCAAGAACTT	ATTTGTACAT	TTTTTTTTC	2160
35	AATAAAACTT	TTCCAATGAC	AAAAAAAA	АААААААА	AAAAAGGGGS	GGGCCGCTCC	2220
	TAGAGGGATC	CCTCCGANGG	NGCCCAATCG	AAAATN			2256
40							
10	(2) INFORM	ATION FOR C	EQ ID NO: 12	n n .			
45	(1)	(A) LEN	HARACTERIST GTH: 829 ba	se pairs			
			E: nucleic ANDEDNESS:				
			OLOGY: line				
50	(xi) SEQUENCE	DESCRIPTION	: SEQ ID NO	: 123:		
	ATGCGCTCCC	TCCTGCTTCT	CAGCGCCTTC	TGCCTCCTGG	AGGCGGCCCT	GGCCGCCGAG	60
55 ·	GTGAAGAAAC	CTGCAGCCGC	AGCAGCTCCT	GGCACTGCGG	AGAAGTTGAG	 CCCCAAGGCG	120
ی ی	GCCACGCTTG	CCGAGCGCAA	GCGGCCTGGC	CTTCAGCTTG	TACCAGGCCA	TGGCCAAGGA	180
	CCAGGCAGTG	GAGAACATCC	TGGTGTCACC	CGTGGTGGTG	GCCTCGTCGC	TGGGGCTCGT	240
60	GTCGCTGGGC	GGCAAGGCGA	CCACGGCGTC	GCAĠGCCAAG	GCAGTGCTGA	GCGCCGAGCA	300

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	GCTGCGCGAC GAGGAGGTGC ACGCCGGCCT GGGCGAGCTG CTGCGCTCAC TCAGCAACTC	360
5	CACGGCGCGC AACGTGACCT GGAAGCTGGG CAGCCGACTG TACGGACCCA GCTCAGTGAG	420
J	CTTCGCTGAT GACTTCGTGC GCAGCAGCAA GCAGCACTAC AACTGCGAGC ACTCCAAGAT	480
	CAACTTCCGC GACAAGCGCA GCGCGCTGCA GTCCATCAAC GAGTGGGCCG CGCAGACCAC	540
10	CGACGCCAAG CTGCCCGAGG TCACCAAGGA CGTGGAGCGC ACGGACGGCG CCCTGTTAGT	600
	CAACGCCATG TTCTTCAAGC CACACTGGGA TGAGAAATTC CACCACAAGA TGGTGGACAA	660
15	CCGTGGCTTC ATGGTGACTC GGTCCTATAC CGTGGGTGTC ATGATGATGC ACCGGACAGG	720
15	CCTCTACAAC TACTACGACG ACGAGAAGGA AAAGCTGCAA ATCGTGGAGA TGCCCCTGGC	780
	CCACAAGCTC TCCAGCCTCA TCATCCTCAT GCCCCATCAC GTGGAGCCT	829
20		
	(2) INFORMATION FOR SEQ ID NO: 124:	
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2223 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 124:	
	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGCCAGGAG GGGGTGGGGA	60
35	CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT	120
	CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG	180
40	CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG	240
10	CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA	300
	AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG AGCGCAGNCG GCCTGGCCTT CAGCTTGTAC	360
45	CAGGCCATGG CCAAGGACCA GGCAGTGGAG AACATCCTGG TGTCACCCGT GGTGGTGGCC	420
	TCGTCGCTGG GGCTCGTGTC GCTGGGCGGC AAGGCGACCA CGGCGTCGCA GGCCAAGGCA	480
50	GTGCTGAGCG CCGAGCAGCT GCGCGACGAG GAGGTGCACG CCGCCCTGGG CGAGCTGCTG	540
	CGCTCACTCA GCAACTCSAC GGCGCGCAAC GTGACCTGGA AGCTGGGCAG CCGACTGTAC	600
	GGACCCAGCT CAGTGAGCTT CGCTGATGAC TTCGTGCGCA CAGCAAGCAG CACTACAACT	660
55	GCGAGCACTC CAAGATCAAC TTCCGCGACA AGCGCACGCG CTGCAGTCCA TCAACGAGTG	720
	GGCCGCGCAG ACCACCGACG GCAAGCTGCC CGAGGTCACC AAGGACGTGG AGCGCACGGA	780
60	CGGCGCCCTG YTAGTCAACG CCATGTTCTT CAAGCCACAC TGGGATGAGA AATTCCACCA	840

	CAAGATGGTG	GACAACCGTG	GCTTCATGGT	GACTCGGTCC	TATACYGTGG	GTGTCATGAT	900
	GATGCACCGG	ACAGGCCTCT	ACAACTACTA	CGACGACGAG	AAGGAAAAGC	TGCAAATCGT	960
5	GGAGATGCCC	CTGGCCCACA	AGCTCTCCAG	CCTCATCATC	CTCATGCCCC	ATCACGTGGA	1020
	GCCTCTCGAG	CGCCTTGAAA	AGCTGCTAAC	CAAAGAGCAG	CTGAAGATCT	GGATGGGGAA	1080
10	GATGCAGAAG	AAGGCTGTTG	CCATCTCCTT	GCCCAAGGGT	GTGGTGGAGG	TGACCCATGA	1140
10	CCTGCAGAAA	CACCTGGCTG	GGCTGGGCCT	GACTGAGGCC	ATTGACAAGA	ACAAGGCCGA	1200
	CTTRTCACGC	ATGTCAGGCA	AGAAGGACCT	GTACCTGGCC	AGCGTGTTCC	ACGCCACCGC	1260
15	CTTTGAGTTG	GACACAGATG	GCAACCCCTT	TGACCAGGAC	ATCTACGGGC	GCGAGGAGCT	1320
	GCGCASCCCA	AGCTGTTCTA	CGCCGACCAC	CCCTTCATCT	TCCTAGTGCG	GGACACCCAA	1380
20	AGCGGCTCCC	TGCTATTCAT	TGGGCGCCTG	GTCCGGCCTA	AGGGTGACAA	GATGCGAGAC	1440
20	GAGTTATAGG	GCCTCAGGGT	GCACACAGGA	TGGCAGGAGG	CATCCAAAGG	CTCCTGAGAC	1500
	ACATGGGTGC	TATTGGGGTT	GGGGGGGAGG	TGAGGTACCA	GCCTTGGATA	CTCCATGGGG	1560
25	TGGGGGTGGA	AAARCAGACC	GGGGTTCCCG	TGTGCCTGAG	CGGACCTTCC	CAGCTAGAAT	1620
	TCACTCCACT	TGGACATGGG	CCCCAGATAC	CATGATGCTG	AGCCCGGAAA	CTCCACATCC	1680
30	TGTGGGACCT	GGCCATAGT	CATTCTGCCT	GCCCTGAAAG	TCCCAGATCA	AGCCTGCCTC	1740
	AATCAGTATT	САТАТТТАТА	GCCAGGTACC	TTCTCACCTG	TGAGACCAAA	TTGAGCTAGG	1800
	GGGTCAGCC	AGCCCTCTTC	TGACACTAAA	ACACCTCAGC	TGCCTCCCCA	GCTCTATCCC	1860
35	AACCTCTCCC	AACTATAAAA	CTAGGTGCTG	CAGCCCCTGG	GACCAGGCAC	CCCCAGAATG	1920
	ACCTGGCCGC	AGTGAGGCGG	ATTGAGAAGG	AGCTCCCAGG	AGGGGCTTCT	GGGCAGACTC	1980
40	TGGTCAAGAA	GCATCGTGTC	TGGCGTTGTG	GGGATGAACT	TTTTGTTTTG	TTTCTTCCTT	2040
.0	TTTTAGTTCT	TCAAAGATAG	GGAGGGAAGG	GGGAACATGA	GCCTTTGTTG	CTATCAATCC	2100
	AAGAACTTAT	TTGTACATTT	TTTTTTCAA	TAAAACTTTT	CCAATGACAA	AAAAAAAAA	2160
45	AAAAAAAA	MWMGGGGSGG	GCCGCTCCTA	GAGGGATCCC	TCCGANGGNG	CCCAATCGAA	2220
	AAT						222

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(2) INFORMATION FOR SEQ ID NO: 125:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 31 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 125:

 $60\,$ Met Lys Lys Gln Ser Lys Arg Cys Leu Trp Lys Pro Pro Gly Ser Leu

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	1				5					10					15	
5	Arg	Arg	Leu	Trp 20	Trp	Met	Arg	Ala	Leu 25	Leu	Ile	Leu ·	Lys	Туг 30	Ile	
	(2)	INFO	ORMA!	rion	FOR	SEQ	ID 1	10: 1	126:					•		
10				(1	A) L: B) T D) T	ENGT YPE : OPOLA	H: 4 ami OGY:	5 am no a lin	ino cid ear	acid		. 10	c .			
15	Met 1		_	Ser										Met	Leu 15	Leu
20	His	Leu	Thr	Ala 20	Ala	Phe	Leu	Gln	Arg 25	Ala	His	Xaa	Ile	Leu 30	Thr	Thr
	Arg	Met	Ser 35	Leu	Gly	Phe	Gln	Ser 40	Pro	His	Leu	Thr	Met 45	`		
25	(2)	INF	ORMA'	NOIT	FOR	SEQ	ID I	NO: :	127 :							
30				(A) L B) T D) T	ENGT YPE : OPOL	H: 3 ami OGY:	9 am no a lin	ino cid ear	acid		: 12	7:			
35	Met 1	His	Asn	Gln	Arg 5	Gln	Val	Phe	Leu	Phe 10	His	Leu	Phe	Ser	Asn 15	Tyr
40				Ile 20 Met				Pro	Gly 25	Thr	Leu	Leu	Ala	. Ala 30	Thr	Tyr
	Cys	Dea	35	riec	1111	171	GIY									
45	(2)	INF		TION												
50				(A) L B) T D) T	ENGT YPE : YPOL	H: Z ami OGY:	23 an ino a : lir	mino acid near	acio): 12	28:			
55	Met 1		Lys	Lys	Phe 5		Leu	Ala	Gln	Val		. Leu	ı Sei	. Leu	Ser 15	Val
	Met	Pro) Ser	Met 20		Val	Thr	•								

	(2)	INF	DRMAT	NOI	FOR	SEQ	ID N	10: 1	.29:							
5			(i) 5	(; (;	A) L B) T D) T	ENGT: YPE : OPOL	H: 1 ami: OGY:	10 au no ao lino	mino cid ear	aci		. 12	n .			
			(xi)													
10	Met 1		Leu	Leu	Cys 5	Leu	Leu	Leu	Val	Pro 10	Leu	Leu	Leu	Ser	Leu 15	Phe
15	Val	Leu	Gly	Leu 20	Phe	Leu	Trp	Phe	Leu 25	Lys	Arg	Glu	Arg	Gln 30	Gĺu	Glu
15	Tyr	Ile	Glu 35	Glu	Lys	Lys	Arg	Val 40	Asp	Ile	Cys	Arg	Glu 45	Thr	Pro	Asn
20	Ile	Cys 50	Pro	His	Ser	Gly	Glu 55	Asn	Thr	Glu	Tyr	Asp 60	Thr	Ile	Pro	His
	Thr 65	Asn	Arg	Thr	Ile	Leu 70	Lys	Glu	Asp	Pro	Ala 75	Asn	Thr	Val	Tyr	Ser 80
25	Thr	Val	Glu	Ile	Pro 85	Lys	Lys	Met	Glu	Asn 90	Pro	His	Ser	Leu	Leu 95	Thr
30	Met	Pro	Asp	Thr 100	Pro	Arg	Leu	Phe	Ala 105	Tyr	Glu	Asn	Val	Ile 110		
	(2)	INF	ORMA'	rion	FOR	SEQ	ID i	NO:	130:							
35				(A) I B) T D) T	CHA LENGT TYPE: TOPOL E DE	TH: 6 ami LOGY:	3 am no a lir	ino cid ear	acid		v. 12	0.			
40			(X1)	SEQ	OEMC	E DE	CKI		W: 5	EQ I	טאז עו.	/. IJ	U:			
	Met 1		Leu	Leu	Phe 5		Tyr	Phe	Tyr	Ser 10	His	Pro	Ala	Pro	Val 15	Pro
45	Ala	Gly	Ala	Thr 20		Lys	Pro	Arg	Туг 25	_	Val	Ile	Thr	Cys 30	-	Pro
	Ala	Ser	Val 35		Ser	Thr	Ser	Phe 40		His	Ser	Pro	Pro 45		. Arg	Cys
50	Leu	Gly 50	/ Arg	Leu	Glu	Gln	Met 55		His	Phe	: Gly	Leu 60		Ser	Gly	•
55 .	(2)	INF	ORMA													
			(i)			E CHA LENG					ds					
60						TYPE TOPO:										
					/											

```
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 131:
     Met Pro Phe Pro Ile Ser Ile Leu Gln Leu Cys Leu Gln Ile Ser Asn
5
     Leu Ser Phe Cys Leu Gln Lys Ile Tyr Lys Ile Pro Phe Val
10
     (2) INFORMATION FOR SEQ ID NO: 132:
             (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 53 amino acids
15
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 132:
     Met Ala Ala Cys Arg Ser Val Lys Gly Leu Val Ala Val Ile Thr
20
     Gly Gly Ala Ser Gly Leu Gly Leu Ala Thr Ala Asp Asp Leu Trp Gly
25
     Arg Glu Pro Leu Cys Phe Trp Thr Cys Pro Thr Arg Val Gly Arg
                                 40
     Pro Lys Pro Arg Ser
          50
30
      (2) INFORMATION FOR SEQ ID NO: 133:
35
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 57 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 133:
40
     Met Leu Leu Val Tyr Asp Leu Tyr Leu Xaa Pro Lys Leu Trp Ala Leu
                                10
      Ala Thr Pro Gln Lys Asn Gly Lys Gly Ala Arg Xaa Gly Asp Gly Thr
45
      Pro Ala Gln Ala Phe Trp Asp Phe Trp Ser His Leu Ile Ser Ala Asp
50
      Pro Gln Thr Trp Glu Arg Ala Ala Pro
           50
55
      (2) INFORMATION FOR SEQ ID NO: 134:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 216 amino acids
                    (B) TYPE: amino acid
60
                    (D) TOPOLOGY: linear
```

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 134:

(x1) SEQUENCE DESCRIPTION: SEQ ID NO: 134:																
. 5	Met 1	Arg	Leu	Ser	Ala 5	Leu	Leu	Ala	Leu	Ala 10	Ser	Lys	Val	Thr	Leu 15	Pro
5	Pro	His	Tyr	Arg 20	Tyr	Gly	Met	Ser	Pro 25	Pro	Gly	Ser	Val	Ala 30	Asp	Lys
10	Arg	Lys	Asn 35	Pro	Pro	Trp	Ile	Arg 40	Arg	Arg	Pro	Val	Val 45	Val	Glu	Pro
	Ile	Ser 50	Asp	Glu	Asp	Trp	Tyr 55	Leu	Phe	Cys	Gly	Asp 60	Thr	Val	Glu	Ile
15	Leu 65	Glu	Gly	Lys	Asp	Ala 70	Gly	Lys	Gln	Gly	Lys 75	Val	Val	Gln	Val	Ile 80
20	Arg	Gln	Arg	Asn	Trp 85	Val	Val	Val	Gly	Gly 90	Leu	Asn	Thr	His	Tyr 95	Arg
	Tyr	Ile	Gly	Lys 100	Thr	Met	Asp	Tyr	Arg 105		Thr	Met	Ile	Pro 110	Ser	Glu
25	•		Leu 115			,		120	-				125			
		130					135					140				
30	145		Ser			150					155					160
35					165	•				170)				175	
				180)				185	5				190	l	Lys Arg
40			195		_			200)	i ne	- 01)		209			9
45	Буз	210		, Dy S	, va		215		-							
	(2)	IN	FORMA	OITA	N FOI	R SE	Q ID	NO:	135	:						
50					(A) (B) (D)	LENC TYPE TOPO	ARAC TH: : am OLOGY ESCR	49 a ino : li	mino acid near	aci l		iO: 1	35 :			
55		t Se 1	r Le	u Ar		n Ly 5	s Se	r Se	r Ph		g Le 0	u Me	t Va	l Me	t Se 1	r Leu 5
60	Th	r Il	e Le		s Le O	u Se	r Ly	s Th		r Va 5	l Le	u Cy	s Le		g Су 0	s Leu

His Ser Leu Lys Leu Thr Trp Arg Asp Gly Ala Arg Cys Ile Asn Ala 40 Glu 5 (2) INFORMATION FOR SEQ ID NO: 136: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 68 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 15 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 136: Met Ser Gly Ser Phe Ile Leu Cys Leu Ala Leu Val Thr Arg Trp Ser 5 20 Pro Gln Ala Ser Ser Val Pro Leu Ala Val Tyr Glu Ser Lys Thr Arg 25 Lys Ser Tyr Arg Ser Gln Arg Asp Arg Asp Gly Lys Asp Arg Ser Gln 40 25 Gly Met Gly Leu Ser Leu Leu Val Glu Thr Arg Lys Leu Leu Ser 55 Ala Asn Gln Gly 30 65 (2) INFORMATION FOR SEQ ID NO: 137: 35 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 52 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 137: Met Cys Phe Arg Phe Phe Leu Phe Cys Ser Arg Ile Leu Leu Lys Leu 45 Phe Phe Leu Leu Phe Pro Ala Ser Ala Phe Pro Leu Ser Thr Arg Ser Ser Leu Ser Val Asn Glu His Val Val Val Ser Pro Arg Ser Thr Val 50 Ser Ile Ser Arg 50 55 (2) INFORMATION FOR SEQ ID NO: 138: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 541 amino acids 60 (B) TYPE: amino acid

(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 138:

5	Met 1	Val	Arg	Thr	Asp 5	Gly	His	Thr	Leu	Ser 10	Glu	Lys	Arg	Asn	Туг 15	Gln
	Val	Thr	Asn	Ser 20	Met	Phe	Gly	Ala	Ser 25	Arg	Lys	Lys	Phe	Val 30	Glu	Gly
10	Val	Asp	Ser 35	Asp	Туг	His	Asp	Glu 40	Asn	Met	Tyr	Tyr	Ser 45	Gln	Ser	Ser
15	Met	Phe 50	Pro	His	Arg	Ser	Glu 55	Lys	Asp	Met	Leu	Ala 60	Ser	Pro	Ser	Thr
	Ser 65	Gly	Gln	Leu	Ser	Gln 70	Phe	Gly	Ala	Ser	Leu 75	Tyr	Gly	Gln	Gln	Ser 80
20	Ala	Leu	Gly	Leu	Pro 85	Met	Arg	Gly	Met	Ser 90	Asn	Asn	Thr	Pro	Gln 95	Leu
	Asn	Arg	Ser	Leu 100	Ser	Gln	Gly	Thr	Gln 105	Leu	Pro	Ser	His	Val 110	Thr	Pro
25	Thr	Thr	Gly 115	Val	Pro	Thr	Met	Ser 120	Leu	His	Thr	Pro	Pro 125	Ser	Pro	Ser
30		130		Leu			135					140				
	145			Gly		150					155					160
35				Gly	165					170					175	
40				Gln 180					185					190		
40			195	Met				200					205			
45		210		Ile			215					220				
	225			Ser		230					235					240
50				Asn	245					250					255	•
~ ~ ~				Gly 260					265					270		
55	· Phe		275					280					285			
60	Lys	Asp 290	Pro	Thr	Ser	Ser	Asn 295	Asp	Asp	Ser	Lys	Ser 300	Asn	Leu	Asn	Thr

	Ser 305	Gly	Lys	Thr	Thr	Ser 310	Ser	Thr	Asp	Gly	Pro 315	Lys	Phe	Pro	Gly	Asp 320
5 .	Lys	Ser	Ser	Thr	Thr 325	Gln	Asn	Asn	Asn	Gln 330	Gln	Lys	Lys	Gly	Ile 335	Gln
	Val	Leu	Pro	Asp 340	Gly	Arg	Val	Thr	Asn 345	Ile	Pro	Gln	Gly	Met 350	Val	Thr
10	Asp	Gln	Phe 355	Gly	Met	Ile	Gly	Leu 360	Leu	Thr	Phe	Ile	Arg 365	Ala	Ala	Glu
15	Thr	Asp 370	Pro	Gly	Met	Val	His 375	Leu	Ala	Leu	Gly	Ser 380	Asp	Leu	Thr	Thr
	Leu 385	Gly	Leu	Asn	Leu	Asn 390	Ser	Pro	Glu	Asn	Leu 395	Tyr	Pro	Lys	Phe	Ala 400
20	Ser	Pro	Trp	Ala	Ser 405	Ser	Pro	Cys	Arg	Pro 410		Asp	Ile	Asp	Phe 415	His
	Val	Pro	Ser	Glu 420	Tyr	Leu	Thr	Asn	Ile 425	His	Ile	Arg	Asp	Lys 430	Leu	Ala
25	Ala	Ile	Lys 435	Leu	Gly	Arg	Tyr	Gly 440	Glu	Asp	Leu	Leu	Phe 445	Tyr	Leu	Tyr
30	Tyr	Met 450	Asn	Gly	Gly	Asp	Val 455	Leu	Gln	Leu	Leu	Ala 460	Ala	Val	Glu	Leu
	Phe 465	Asn	Arg	Asp	Trp	Arg 470	Tyr	His	Lys	Glu	Glu 475	Arg	Val	Trp	Ile	Thr 480
35	Arg	Ala	Pro	Gly	Met 485	Glu	Pro	Thr	Met	Lys 490	Thr	Asn	Thr	Tyr	Glu 495	Arg
	Gly	Thr	Tyr	туr 500	Phe	Phe	Asp	Cys	Leu 505	Asn	Trp	Arg	Lys	Val 510	Ala	Lys
40	Glu	Phe	His 515	Leu	Glu	Tyr	Asp	Lys 520	Leu	Glu	Glu	Arg	Pro 525	His	Leu	Pro
45	Ser	Thr 530	Phe	Asn	Tyr	Asn	Pro 535	Ala	Gln	Gln	Ala	Phe 540	Xaa			
	(2)	INFO	RMAT	CION	FOR	SEQ	ID N	10: 1	.39:							
50			(i) S	(A) L B) T	ENGT: YPE:	H: 5 ami	ERIST 8 am no a lin	ino . cid		s					
55			(xi)					PTIO		EQ II	D NO	: 13	9 :			
	Met 1	Ile	Cys	Pro	Gln 5	Cys	Pro	Leu	Ser	Leu 10	Leu	Cys	Leu	Ile	Ser 15	·Ser
60	Leu	Cys	Ser	Leu 20	Val	Ile	Gln	Ile	Ser 25	Leu	Lys	Thr	Ile	Arg 30	Asp	Ile

	Thr	Leu	Leu 35	Asn	Met	Val	Gly	Ile 40	Lys	Phe	Ser	Ile	Ser 45	Leu	Ser	Asn
5	Lys	Ile 50	Asn	Ile	Asn	Ser	Arg 55	Thr	Trp	Xaa						
10	(2)	INFO	ORMA!	NOI	FOR	SEQ	ID I	NO: 1	140:							
15			(i) :	(A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami OGY:	02 a no a lin	mino cid ear	aci		: 14	0 :		•	
20	Met 1	Thr	Leu	Arg	Pro 5	Ser	Leu	Leu	Pro	Leu 10	His	Leu	Leu	Leu	Leu 15	Leu
20	Leu	Leu	Ser	Ala 20	Ala	Val	Cys	Arg	Ala 25	Glu	Ala	Gly	Leu	Glu 30	Thr	Glu
25	Ser	Pro	Val 35	Arg	Thr	Leu	Gln	Val 40	Glu	Thr	Leu	Val	Glu 45	Pro	Pro	Glu
	Pro	Cys 50	Ala	Glu	Pro	Ala	Ala 55	Phe	Gly	Asp	Thr	Leu 60	His	Ile	His	Tyr
30	Thr 65	Gly	Ser	Leu	Val	Asp 70	Gly	Arg	Ile	Ile	Asp 75	Thr	Ser	Leu	Thr	Arg 80
35	Asp	Pro	Leu	Val	Ile 85	Glu	Leu	Gly	Gln	Lys 90	Gln	Val	Ile	Pro	Gly 95	Leu
))	Glu	Gln	Ser	Leu 100	Leu	Asp	Met	Суѕ	Val 105	Gly	Glu	Lys	Arg	Arg 110	Ala	Ile
40	Ile	Pro	Ser 115	His	Leu	Ala	Tyr	Gly 120	Lys	Arg	Gly	Phe	Pro 125	Pro	Ser	Val
	Pro	Ala 130	Asp	Ala	Val	Val	Gln 135	Tyr	Asp	Val	Glu	Leu 140	Ile	Ala	Leu	Ile
45	Arg 145	Ala	Asn	Tyr	Trp	Leu 150	Lys	Leu	Val	Lys	Gly 155	Ile	Leu	Pro	Leu	Val
	Gly	Met	Ala	Met	Val 165	Pro	Ala	Leu	Leu	Gly 170	Leu	Ile	Gly	Tyr	His 175	Leu
50	Tyr	Arg	Lys	Ala 180	Asn	Arg	Pro	Lys	Val 185	Ser	Lys	Lys	Lys	Leu 190	Lys	Glu
55	Glu	Lys	Arg 195	Asn	Lys	Ser	Lys	Lys 200	Lys	Xaa						

(2) INFORMATION FOR SEQ ID NO: 141:

	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 217 amino acids(B) TYPE: amino acid(D) TOPOLOGY: linear															
5 .	5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 141:															
	Met 1	Phe	Leu	Arg	Leu 5	Тут	Leu	Ile	Ala	Arg 10	Val	Met	Leu	Leu	His 15	Ser
10	Lys	Leu	Phe	Thr 20	Asp	Ala	Ser	Ser	Arg 25	Ser	Ile	Gly	Ala	Leu 30	Asn	Lys
15	Ile	Asn	Phe 35	Asn	Thr	Arg	Phe	Val 40	Met	Lys	Thr	Leu	Met 45	Thr	Ile	Cys
	Pro	Gly 50	Thr	Val	Leu	Leu	Val 55	Phe	Ser	Ile	Ser	Leu 60	Trp	Ile	Ile	Ala
20	Ala 65	Trp	Thr	Val	Arg	Val 70	Cys	Glu	Ser	Pro	Glu 75	Ser	Pro	Ala	Gln	Pro 80
	Ser	Gly	Ser	Ser	Leu 85	Pro	Ala	Trp	Tyr	His 90	Asp	Gln	Gln	Asp	Val 95	Thr
25	Ser	Asn	Phe	Leu 100	Gly	Ala	Met	Trp	Leu 105	Ile	Ser	Ile	Thr	Phe 110	Leu	Ser
30	Ile	Gly	Туr 115	Gly	Asp	Met	Val	Pro 120	His	Thr	Tyr	Cys	Gly 125	Lys	Gly	Val
	Cys	Leu 130	Leu	Thr	Gly	Ile	Met 135	Gly	Ala	Gly	Cys	Thr 140	Ala	Leu	Val	Val
35	Ala 145	Val	Val	Ala	Arg	Lys 150	Leu	Glu	Leu	Thr	Lys 155	Ala	Glu	Lys	His	Val 160
	His	Asn	Phe	Met	Met 165	Asp	Thr	Gln	Leu	Thr 170	Lys	Arg	Ile	Lys	Asn 175	Ala
40	Ala	Ala	Asn	Val 180	Leu	Arg	Glu	Thr	Trp 185	Leu	Ile	Tyr	Lys	His 190	Thr	Lys
45	Leu	Leu	Lys 195	Lys	Ile	Asp	His	Ala 200	Lys	Val	Arg	Lys	His 205	Gln	Arg	Lys
	Phe	Leu 210	Pro	Ser	Tyr	Pro	Pro 215	Val	Xaa							
50	(2)	TATE	N 10 14 N 10	TON	505											
	(2)		ORMAT													
55			(i) :	(A) L B) T D) T	ENGT YPE : OPOL	H: 1 ami OGY:	02 a no a lin	mino cid ear	aci						
											D NO					
60	Met 1	ser	Asn	Thr	Thr 5	Val	Pro	Asn	Ala	Pro 10	Gln	Ala	Asn	Ser	Asp 15	Ser

	Met	Val	Gly	Tyr 20	Val	Leu	Gly	Pro	Phe 25	Phe	Leu	Ile	Thr	Leu 30	Val	Gly
5	Val	Val	Val 35	Ala	Val	Val	Met	Tyr 40	Val	Gln	Lys	Lys	Lys 45	Arg	Val	Asp
10	Arg	Leu 50	Arg	His	His	Leu	Leu 55	Pro	Met	Tyr	Ser	Туr 60	Asp	Pro	Ala	Glu
	Glu 65	Leu	His	Glu	Ala	Glu 70	Gln	Glu	Leu	Leu	Ser 75	Asp	Met	Gly	Asp	Pro 80
15	Lys	Val	Val	His	Gly 85	Trp	Gln	Ser	Gly	Tyr 90	Gln	His	Lys	Arg	Met 95	Pro
	Leu	Leu	Asp	Val 100	Lys	Thr										
20	(2)	INFO	ORMA	rion	FOR	SEQ	I di	NO: í	143:							
25	(2) INFORMATION FOR SEQ ID NO: 143: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 112 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (Xi) SEQUENCE DESCRIPTION: SEQ ID NO: 143:															
30	Met 1	Arg	Glu	Cys	Gln 5	Glu	Glu	Ser	Phe	Trp 10	Lys	Arg	Ala	Leu	Pro 15	Phe
35	Ser	Leu	Val	Ser 20	Met	Leu	Val	Thr	Gln 25	Gly	Leu	Val	Tyr	Gln 30	Gly	Tyr
	Leu	Ala	Ala 35	Asn	Ser	Arg	Phe	Gly 40	Ser	Leu	Pro	Lys	Val 45	Ala	Leu	Ala
40	Gly	Leu 50	Leu	Gly	Phe	Gly	Leu 55	Gly	Lys	Val	Ser	Tyr 60	Ile	Gly	Val	Cys
	Gln 65	Ser	Lys	Phe	His	Phe 70	Phe	Glu	Asp	Gln	Leu 75	Arg	Gly	Ala	Gly	Phe 80
45	Gly	Pro	Gln	His	Asn 85	Arg	His	Cys	Leu	Leu 90	Thr	Cys	Glu	Glu	Cys 95	Lys
50	Ile	Lys	His	Gly 100	Leu	Ser	Glu	Lys	Gly 105	Asp	Ser	Gln	Pro	Ser 110	Ala	Ser
55 -	(2)	INF	orma'	TION	FOR	SEO	ID:	NO:	144:		••					
	(2) INFORMATION FOR SEQ ID NO: 144: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 amino acids															

(B) TYPE: amino acid

```
(D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 144:
      Met Lys Asn Asp Arg Asn Gln Gly Phe Ser Leu Leu Gln Leu Ile Asp
 5
      Trp Asn Lys Pro
10
      (2) INFORMATION FOR SEQ ID NO: 145:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 30 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 145:
20
      Met Gly Thr Gln Pro Pro Val Val Ala Gly Phe Thr Ile Pro Met Leu
     Gly Tyr Thr Val Arg Val Leu Thr Phe His Leu Ser Cys Ser
                   20
25
      (2) INFORMATION FOR SEQ ID NO: 146:
30
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 99 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 146:
35
     Met Lys Ile Pro Val Leu Pro Ala Val Val Leu Leu Ser Leu Leu Val
                                           10
      Leu His Ser Ala Gln Gly Ala Thr Leu Gly Gly Pro Glu Glu Glu Ser
40
                                       25
     Thr Ile Glu Asn Tyr Ala Ser Arg Pro Glu Ala Phe Asn Thr Pro Phe
45
     Leu Asn Ile Asp Lys Leu Arg Ser Ala Phe Lys Ala Asp Glu Phe Leu
          50
     Asn Trp His Ala Leu Phe Glu Ser Ile Lys Arg Lys Leu Pro Phe Leu
50
     Asn Trp Asp Ala Phe Pro Lys Leu Lys Gly Leu Arg Ser Ala Thr Pro
                                           90
     Asp Ala Gln
55
```

(2) INFORMATION FOR SEQ ID NO: 147:

```
(i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 8 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
5
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 147:
     Met Val Trp Gly Leu Leu Gly
10
      (2) INFORMATION FOR SEQ ID NO: 148:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 39 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 148:
20
      Met Leu Pro Leu Leu Ser Leu Leu Phe Leu Phe Phe Ser Thr Val Ser
      Ser Phe Cys Gly Met Pro Leu Arg Ala His Thr Arg Ala Xaa Ala His
25
      Thr Arg Thr Phe Ala Ser Arg
               35
30
      (2) INFORMATION FOR SEQ ID NO: 149:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 131 amino acids
35
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 149:
      Met Ile Cys Glu Thr Lys Ala Arg Lys Ser Ser Gly Gln Pro Gly Arg
40
                        5
      Leu Pro Pro Pro Thr Leu Ala Pro Pro Gln Pro Pro Leu Pro Glu Thr
45
      Ile Glu Arg Pro Val Gly Thr Gly Ala Met Val Ala Arg Ser Ser Asp
               35
                                   40
                                                        45
      Leu Pro Tyr Leu Ile Val Gly Val Val Leu Gly Ser Ile Val Leu Ile
                               55
50
      Ile Val Thr Phe Ile Pro Phe Cys Leu Trp Arg Ala Trp Ser Lys Gln
                           70
                                                75
      Lys His Thr Thr Asp Leu Gly Phe Pro Arg Ser Ala Leu Pro Pro Ser
55
                                            90
      Cys Pro Tyr Thr Met Val Pro Leu Gly Gly Leu Pro Gly His Gln Ala
                  100
                                       105
                                                           110
60
      Val Asp Ser Pro Thr Ser Val Ala Ser Val Asp Gly Pro Val Leu Met
```

```
115
                                  120
                                                      125
     Gly Ser Thr
         130
 5
      (2) INFORMATION FOR SEQ ID NO: 150:
10
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 32 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 150:
15
     Met Gly Ala Pro Ser Leu Thr Met Leu Leu Leu Leu Lys Val Gln Pro
      Arg Arg Thr Gln Ala Phe Asp Ala His Trp Val Gly Leu Pro Leu Leu
20
                                       25
25
      (2) INFORMATION FOR SEQ ID NO: 151:
             (i) SEQUENCE CHARACTERISTICS:
30
                    (A) LENGTH: 14 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 151:
35
      Met Cys Leu Ile Phe Leu Leu Leu Leu Leu Ser Phe Ser
                        5
        1
                                          10
40
      (2) INFORMATION FOR SEQ ID NO: 152:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 8 amino acids
                     (B) TYPE: amino acid
45
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 152:
      His Pro His Gln Asp Ser Gln Pro
       1
                       5
50
      (2) INFORMATION FOR SEQ ID NO: 153:
55
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 68 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 153:
60
```

	Met 1	Asn	Thr	Ser	Tyr 5	Ile	Leu	Arg	Leu	Thr 10	Val	Val	Val	Ser	Val ' 15	Val
5	Ile	туr	Leu	Ala 20	Ile	His	Pro	Leu	Leu 25	Ser	Phe	Ser	Leu	Glu 30	Ser	Pro
	Leu	Leu	Val 35	Pro	Trp	Arg	Asp	Cys 40	Суз	Gln	Asn	Ile	Trp 45	Lys	Ser	Gly
10	Ser	Val 50	Trp	Tyr	Lys	Arg	Trp 55	Thr	Leu	Pro	His	Met 60	Glu	Val	Cys	Cys
15	Gln 65	Asp	Leu	His												
	(2)	INF	ORMA	rion	FOR	SEQ	ID 1	۷O: ۱	154:							
20			(i) .	(A) L B) T		H: 2 ami	6 am no a			s					
25			(xi)						N: S	EQ I	D NO	: 154	4 :			
	Met 1		Lys	Ile	Phe 5	Lys	Glu	Trp	Glu	Asn 10	Leu	Asn	Leu	Ile	Leu 15	Thr
30	Ser	Ile	Arg	Ile 20	Leu	Glu	Arg	Gln	Asn 25	Met						
35	(2)	INF	ORMA'	SEQU	ENCE	СНА	RACT	ERIS	TICS							
				(В) Т	ENGT YPE : OPOL	ami	no a		aci	.ds					
40			(xi)						N: S	EQ I	D NO	: 15	5 :			
	Met 1		Cys	Glu	Val 5		Asn	Gly	Ser	Ser 10		Arg	Asp	Glu	Cys 15	Ile
45	Thr	Asn	Leu	Leu 20		Phe	Gly	Phe	Leu 25		. Ser	Cys	Ser	qzA 06	Asn	Ser
50	Phe	Arg	Arg 35		Leu	Asp	Ala	Leu 40		His	Glu	Leu	Pro 45	Val	Leu	Ala
50	Pro	Glr 50		Glu	Gly	Tyr	Asp 55		ı Leu	Gln	Thr	Asp 60		' Asn	Arg	Ser
55	Ser 65		s Ser	Arg	, Leu	Gly 70		ı Ile	e Glu	Ala	Asp 75		Glu	. Ser	Gln	Glu 80
	Asp) Ile	e Il∈	e Arg	Asn 85		Ala	a Arg	g His	: L et		a Gln	ı Val	Gly	qaA 95	
60	Met	Asp	Arg	, Ser	: Ile	Pro	Pro	Glv	/ Lei	ı Val	L Asr	ı Glv	, Lei	ı Ala	Leu	Gln

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				100					105					110			
5	Leu	Arg	Asn 115	Thr	Ser	Arg	Ser	Glu 120	Glu	Asp	Arg	Asn	Arg 125	Asp	Leu	Ala	
3	Thr	Ala 130	Leu	Glu	Gln	Leu	Leu 135	Gln	Ala	Tyr	Pro	Arg 140	Asp	Met	Glu	Lys	
10	Glu 145	Lys	Thr	Met	Leu	Val 150	Leu	Ala	Leu	Leu	Leu 155	Ala	Lys	Lys	Val	Ala 160	
	Ser	His	Thr	Pro	Ser 165	Leu	Leu	Arg	Asp	Val 170	Phe	His	Thr	Thr	Val 175	Asn	
15	Phe	Ile	Asn	Gln 180	Asn	Leu	Arg	Thr	Тут 185	Val	Arg	Ser	Leu	Ala 190	Arg	Asn	
20	Gly	Met	Asp 195														
	(2)	INFO	ORMA?	гіои	FOR	SEQ	ID I	NO: :	156:								
25			(i)	(A) L B) T	ENGT YPE :	H: 9 ami		ino cid	: acid	s						
30			(xi)	SEQ						EQ I	D NO	: 15	6:				
	Met 1	Ser	Leu	Ser	Leu 5	Val	Ser	Val	Ser	Val 10	Gly	Pro	Ser	Thr	Leu 15	Ala	
35	Cys	Ser	Phe	Leu 20	Arg	Pro	Lys	Ala	Arg 25		Ser	Lys	Arg	Ser 30	Pro	Arg	
	Asn	Tyr	Thr 35	Asp	Ser	Thr	Ser	Pro 40	Gly	Gly	Pro	Arg	Ala 45	Pro	Arg	Gly	
40	Gly	Ala 50	Trp	Arg	Leu	Ser	Ser 55	Gln	Gln	Asn	Ser	Ser 60		Lys	Gly	Val	
45	Ala 65	Val	Ala	Lys	Ala	Ser 70	Tyr	Arg	Pro	Val	Leu 75	Cys	Phe	Leu	Pro	Gly 80	
	Pro	Trp	Ser	Ser	Xaa 85	Pro	Xaa	Ala	Phe	Leu 90	Ile						
50	(2)	INF	ORMA	NOIT	FOR	SEQ	ID	NO:	157 :								
55				(A) I B) T (D) T	ENGT YPE: OPOL	H: 3 am: OGY	31 an ino a : lir	mino acid near	ació): 1 <u>5</u>	5 7 :				
60	Met 1		Thr	Leu	Ser 5		Glu	Cys	Ser	Gly		Ala	. Thr	Leu	Gly 15	, Leu	

Cys Leu Val Val Pro Trp Asn Ser Ser Gly Leu Ser Gln Pro Pro 20 25 5 (2) INFORMATION FOR SEQ ID NO: 158: (i) SEQUENCE CHARACTERISTICS: 10 (A) LENGTH: 91 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 158: 15 Met Lys Phe Leu Ala Val Leu Val Leu Leu Gly Val Ser Ile Phe Leu 10 Val Ser Ala Gln Asn Pro Thr Thr Ala Ala Pro Ala Asp Thr Tyr Pro 25 20 Ala Thr Gly Pro Ala Asp Asp Glu Ala Pro Asp Ala Glu Thr Thr Ala Ala Ala Thr Thr Ala Thr Thr Ala Ala Pro Thr Thr Ala Thr Thr Ala 25 Ala Ser Thr Thr Ala Arg Lys Asp Ile Pro Val Leu Pro Lys Trp Val 30 Gly Asp Leu Pro Asn Gly Arg Val Cys Pro Xaa 85 35 (2) INFORMATION FOR SEQ ID NO: 159: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 89 amino acids (B) TYPE: amino acid 40 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 159: Met Ile Ile Ser Leu Phe Ile Tyr Ile Phe Leu Thr Cys Ser Asn Thr 5 45 Ser Pro Ser Tyr Gln Gly Thr Gln Leu Gly Leu Gly Leu Pro Ser Ala Gln Trp Trp Pro Leu Thr Gly Arg Arg Met Gln Cys Cys Arg Leu Phe 50 35 40 Cys Phe Leu Leu Gln Asn Cys Leu Phe Pro Phe Pro Leu His Leu Ile 55 Gln His Asp Pro Cys Glu Leu Val Leu Thr Ile Ser Trp Asp Trp Ala 55 75 70 Glu Ala Gly Ala Ser Leu Tyr Ser Pro 85 60

	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	10:	160:							
5			(i)	(A) L B) T	ENGT YPE :	H: 1 ami	74 a no a	mino cid		ds					
10			(xi)	SEQ				lin PTIO		EQ I	D NO	: 16	0 :			
10	Met 1	Ser	Ser	Ala	Ala 5	Ala	Asp	His	Trp	Ala 10	Trp	Leu	Leu	Val	Leu 15	Ser
15	Phe	Val	Phe	Gly 20	Суѕ	Asn	Val	Leu	Arg 25	Ile	Leu	Leu	Pro	Ser 30	Phe	Ser
	Ser	Phe	Met 35	Ser	Arg	Val	Leu	Gln 40	Lys	Asp	Ala	Glu	Gln 45	Glu	Ser	Gln
20	Met	Arg 50	Ala	Glu	Ile	Gln	Asp 55	Met	Lys	Gln	Glu	Leu 60	Ser	Thr	Val	Asn
25	Met 65	Met	Asp	Glu	Phe	Ala 70	Arg	Туr	Ala	Arg	Leu 75	Glu	Arg	Lys	Ile	Asn 80
	Lys	Met	Thr	Asp	Lys 85	Leu	Lys	Thr	His	Val 90	Lys	Ala	Arg	Thr	Ala 95	Gln
30	Leu	Ala	Lys	Ile 100	Lys	Trp	Val	Ile	Ser 105	Val	Ala	Phe	Tyr	Val 110	Leu	Gln
	Ala	Ala	Leu 115	Met	Ile	Ser	Leu	Ile 120	Trp	Lys	Tyr	Tyr	Ser 125	Val	Pro	Val
35	Ala	Val 130	Val	Pro	Ser	Lys	Trp 135	Ile	Thr	Pro	Leu	Asp 140	Arg	Leu	Val	Ala
40	Phe 145	Pro	Thr	Arg	Val	Ala 150	Gly	Gly	Val	Gly	Ile 155	Thr	Cys	Trp	Ile	Leu 160
40	Val	Cys	Asn	Lys	Val 165	Val	Ala	Ile	Val	Leu 170	His	Pro	Phe	Ser		
45																
	(2)		ORMAT													
50			(i) : (xi)	(A) L B) T D) T	ENGT: YPE: OPOL	H: 4 ami OGY:	5 am no a lin	ino d cid ear	acid		: 16	1:			
55	Met 1		Lys											Leu	Leu 15	Leu
	Leu	Val	Gln	Cys 20	Glu	Asn	Cys	Cys	Arg 25	Lys	Asn	Met	Leu	Tyr 30	Asn	Ile
60	Phe	Leu	Asn	Ile	His	Asn	Ile	His	Lys	Phe	Ser	Asn	His			

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35 40 45 5 (2) INFORMATION FOR SEQ ID NO: 162: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 162: Met Val Ala Ser Thr Leu Val Thr Asn Leu Phe Gly Val Ala Phe Ala 1 10 15 Thr Thr Ala Ala Thr Arg Ala 20 20 (2) INFORMATION FOR SEQ ID NO: 163: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 70 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 163: Met Leu Met Ala Pro Val Val Cys Leu Ser Phe Ser Pro Cys Pro Ala 30 1 10 Asp Thr Ser Leu Thr Gly Asp Gly Leu Lys Ala Gly Leu Glu Arg Gly 25 35 Xaa Ala Leu Val Thr Leu Phe Asp Ser Val Thr His Phe Leu Ala His 40 Thr Leu Phe Glu Leu Leu Asp Phe Gln Leu Ala Phe Leu Arg Ser Gly 40 Lys Gln Thr Ala Pro His 45 (2) INFORMATION FOR SEQ ID NO: 164: (i) SEQUENCE CHARACTERISTICS: · (A) LENGTH: 323 amino acids 50 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 164: Met Leu Leu Leu Leu Leu Gly Ser Gly Gln Gly Pro Gln Gln 55 Val Gly Ala Gly Gln Thr Phe Glu Tyr Leu Lys Arg Glu His Ser Leu 20 25

Ser Lys Pro Tyr Gln Gly Val Gly Thr Gly Ser Ser Ser Leu Trp Asn

			35					40					45			
5	Leu	Met 50	Gly	Asn	Ala	Met	Val 55	Met	Thr	Gln	Tyr	Ile 60	Arg	Leu	Thr	Pro
3	Asp 65	Met	Gln	Ser	Lys	Gln 70	Gly	Ala	Leu	Trp	Asn 75	Arg	Val	Pro	Cys	Phe 80
10	Leu	Arg	Asp	Trp	Glu 85	Leu	Gln	Val	His	Phe 90	Lys	Ile	His	Gly	Gln 95	Gly
	Lys	Lys	Asn	Leu 100	His	Gly	Asp	Gly	Leu 105	Ala	Ile	Trp	Tyr	Thr 110	Arg	Asn
15	Arg	Met	Gln 115	Pro	Gly	Pro	Val	Phe 120	Gly	Asn	Met	Asp	Lys 125	Phe	Val	Gly
20	Leu	Gly 130	Val	Phe	Val	Asp	Thr 135	Tyr	Pro	Asn	Glu	Glu 140	Lys	Gln	Gln	Glu
	Arg 145	Val	Phe	Pro	Tyr	Ile 150	Ser	Ala	Met	Val	Asn 155	Asn	Gly	Ser	Leu	Ser 160
25	Tyr	Asp	His	Glu	Arg 165	Asp	Gly	Arg	Pro	Thr 170	Glu	Leu	Gly	Gly	Cys 175	Thr
	Ala	Ile	Val	Arg 180	Asn	Leu	His	Tyr	Asp 185	Thr	Phe	Leu	Val	Ile 190	Arg	Туг
30	Val	Lys	Arg 195	His	Leu	Thr	Ile	Met 200	Met	Asp	Ile	Asp	Gly 205	Lys	His	Glu
35	Trp	Arg 210	Asp	Cys	Ile	Glu	Val 215	Pro	Gly	Val	Arg	Leu 220	Pro	Arg	Gly	Туг
	Tyr 225	Phe	Gly	Thr	Ser	Ser 230	Ile	Thr	Gly	Asp	Leu 235	Ser	Asp	Asn	His	Asp 240
40	Val	Ile	Ser	Leu	Lys 245	Leu	Phe	Glu	Leu	Thr 250	Val	Glu	Arg	Thr	Pro 255	Glu
	Glu	Glu	Lys	Leu 260	His	Arg	Asp	Val	Phe 265	Leu	Pro	Ser	Val	Asp 270	Asn	Met
45	Lys	Leu	Pro 275	Glu	Met	Thr	Ala	Pro 280	Leu	Pro	Pro	Leu	Ser 285	Gly	Leu	Ala
50	Leu	Phe 290	Leu	Ile	Val	Phe	Phe 295	Ser	Leu	Val	Phe	Ser 300	Val	Phe	Ala	Ile
	Val 305	Ile	Gly	Ile	Ile	Leu 310	Tyr	Asn	Lys	Trp	Gln 315		Gln	Ser	Arg	Ly:
55	Arg	Phe	Tyr													

			(1)		A) L	ENGT	H: 3	21 au	mino	aci	ds					
								no a lin								
5			(xi)							EQ II	OM C	: 169	5:			
	Met 1	Pro	Ser	Glu	Туг 5	Thr	Tyr	Val	Lys	Leu 10	Arg	Ser	Asp	Cys	Ser 15	Arg
10	Pro	Ser	Leu	Gln 20	Trp	Tyr	Thr	Arg	Ala 25	Gln	Ser	Lys	Met	Arg 30	Arg	Pro
15	Ser	Leu	Leu 35	Leu	Lys	Asp	Ile	Leu 40	Lys	Cys	Thr	Leu	Leu 45	Val	Phe	Gly
10	Val	Trp 50	Ile	Leu	Тут	Ile	Leu 55	Lys	Leu	Asn	Tyr	Thr 60	Thr	Glu	Glu	Cys
20	Asp 65	Met	Lys	Lys	Met	His 70	Tyr	Val	Asp	Pro	Asp 75	His	Val	Lys	Arg	Ala 80
	Gln	Lys	Tyr	Ala	Gln 85	Gln	Val	Leu	Gln	Lys 90	Glu	Cys	Arg	Pro	Lys 95	Ph∈
25	Ala	Lys	Thr	Ser 100	Met	Ala	Leu	Leu	Phe 105	Glu	His	Arg	Tyr	Ser 110	Val	Asp
30	Leu	Leu	Pro 115	Phe	Val	Gln	Lys	Xaa 120	Pro	Lys	Asp	Ser	Glu 125	Ala	Glu	Ser
	Lys	Tyr 130	Asp	Pro	Pro	Phe	Gly 135	Phe	Arg	Lys	Phe	Ser 140	Ser	Lys	Val	Glr
35	Thr 145	Leu	Leu	Glu	Leu	Leu 150	Pro	Glu	His	qaA	Leu 155	Pro	Glu	His	Leu	Lys 160
	Ala	Lys	Thr	Cys	Arg 165	Arg	Суѕ	Val	Val	Ile 170	Gly	Ser	Gly	Gly	Ile 175	Let
40	His	Gly	Leu	Glu 180	Leu	Gly	His	Thr	Leu 185	Asn	Gln	Phe	Asp	Val 190	Val	Ile
45	Arg	Leu	Asn 195	Ser	Ala	Pro	Val	Glu 200	Gly	Tyr	Ser	Glu	His 205	Val	Gly	Ası
	Lys	Thr 210	Thr	Ile	Arg	Met	Thr 215	Tyr	Pro	Glu	Gly	Ala 220	Pro	Leu	Ser	Ası
50	Leu 225	Glu	Tyr	Tyr	Ser	Asn 230	Asp	Leu	Phe	Val	Ala 235	Val	Leu	Phe	Lys	Se:
	Val	Asp	Phe	Asn	Trp 245	Leu	Gln	Ala	Met	Val 250	Lys	Lys	Glu	Thr	Leu 255	Pro
55	Phe	Trp	Val	Arg 260	Leu	Phe	Phe	Trp	Lys 265	Gln	Val	Ala	Glu	Lys 270	Ile	Pro
60	Leu	Gln	Pro 275	Lys	His	Phe	Arg	Ile 280	Leu	Asn	Pro	Val	Ile 285	Ile	Lys	Gl

```
Thr Ala Phe Xaa His Pro Ser Val Leu Arg Ala Ser Val Lys Val Leu
      Gly Ala Glu Ile Arg Thr Ser Pro Gln Ser Val Ser Leu Pro Leu Ser
 5
                         310
      Xaa
10
      (2) INFORMATION FOR SEQ ID NO: 166:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 31 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 166:
20
      Met Thr Leu Asp Val Gln Thr Val Val Val Phe Ala Val Ile Val Val
      Leu Leu Leu Val Asn Val Ile Leu Met Phe Phe Leu Gly Thr Arg
                   20
                                       25
25
      (2) INFORMATION FOR SEQ ID NO: 167:
30
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 72 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 167:
35
      Met Leu Pro Leu Leu Phe Cys Ala Phe Cys Leu His Lys Leu Gly Pro
                                           10
      Leu Leu Phe Leu Tyr Asp Val Leu Met Xaa His Glu Ala Val Met Arg
40
                                       25
      Thr His Gln Ile Gln Leu Pro Asp Pro Glu Phe Pro Ser Gln Gln Asn
45
      Gln Val Leu Asn Lys Thr Leu Phe Asn Lys Leu Lys Lys Lys Lys
           50
      Lys Lys Lys Xaa Xaa Xaa Lys Lys
                           70
50
      (2) INFORMATION FOR SEQ ID NO: 168:
55
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 282 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 168:
60
```

٠	Met 1	Ala	Ser	Arg	Gly 5	Arg	Arg	Pro	Glu	His 10	Gly	Gly	Pro	Pro	Glu 15	Leu
5	Phe	Туг	Asp	Glu 20	Thr	Glu	Ala	Arg	Lys 25	Tyr	Val	Arg	Asn	Ser 30	Arg	Met
	Ile	Asp	Ile 35	Gln	Thr	Arg	Met	Ala 40	Gly	Arg	Ala	Leu	Glu 45	Leu	Leu	Tyr
10	Leu	Pro 50	Glu	Asn	Lys	Pro	Cys 55	Tyr	Leu	Leu	Asp	Ile 60	Gly	Cys	Gly	Thr
15	Gly 65	Leu	Ser	Gly	Ser	Tyr 70	Leu	Ser	Asp	Glu	Gly 75	His	Tyr	Trp	Val	Gly 80
	Leu	Asp	Ile	Ser	Pro 85	Ala	Met	Leu	Asp	Glu 90	Ala	Val	Asp	Arg	Glu 95	Ile
20	Glu	Gly	Asp	Leu 100	Leu	Leu	Gly	Asp	Met 105	Gly	Gln	Gly	Ile	Pro 110	Phe	Lys
	Pro	Gly	Thr 115	Phe	Asp	Gly	Cys	Ile 120	Ser	Ile	Ser	Ala	Val 125	Gln	Trp	Leu
25	Cys	Asn 130	Ala	Asn	Lys	Lys	Ser 135	Glu	Asn	Pro	Ala	Lys 140	Arg	Leu	Tyr	Cys
30	Phe 145	Phe	Ala	Ser	Leu	Phe 150	Ser	Val	Leu	Val	Arg 155	Gly	Ser	Arg	Ala	Val 160
	Leu	Gln	Leu	Tyr	Pro 165	Glu	Asn	Ser	Glu	Gln 170	Leu	Glu	Leu	Ile	Thr 175	Thr
35	Gln	Ala	Thr	Lys 180	Ala	Gly	Phe	Ser	Gly 185	Gly	Met	Val	Val	Asp 190	Tyr	Pro
	Asn	Ser	Ala 195	Lys	Ala	Lys	Lys	Phe 200	Tyr	Leu	Cys	Leu	Phe 205	Ser	Gly	Pro
40	Ser	Thr 210	Phe	Ile	Pro	Glu	Gly 215	Leu	Ser	Glu	Asn	Gln 220	Asp	Glu	Val	Glu
45	Pro 225	Arg	Glu	Ser	Val	Phe 230		Asn	Glu	Arg	Phe 235		Leu	Arg	Met	Ser 240
	Arg	Arg	Gly	Met	Val 245	Arg	Lys	Ser	Arg	Ala 250		Val	Leu	Glu	Lys 255	
50	Glu	Arg	His	Arg 260		Gln	Gly	Arg	Glu 265		Arg	Pro	Asp	Thr 270		Tyr
	Thr	Gly	Arg 275	Lys	Arg	Lys	Pro	Arg 280		Xaa	•					
55	•															

(2) INFORMATION FOR SEQ ID NO: 169:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 23 amino acids

			(xi)		D) T	OPOL	ami : CGY : SCRI	lin	ear	FO T	n NO	. 16	a .			
5		Leu	Gly							Tyr				Ser		Lys
	1 Leu	Met	Val			Ser	Thr			10				•	15	
10				20												
	(2)	INF	ORMA:	rion	FOR	SEQ	ID	NO: 1	170:							
15			(i) :	(A) L B) T D) T	ENGT YPE : OPOL	H: 3 ami OGY:	28 a no a lin	mino cid ear	aci						
20			(xi)	SEQ	UENC:	E DE	SCRI	PTIO	N: S	EQ I	D NO	: 17	0 :			
	Met 1	Trp	Arg	Pro	Ser 5	Val	Leu	Leu	Leu	Leu 10	Leu	Leu	Leu	Arg	His 15	Gly
25	Ala	Gln	Gly	Lys 20	Pro	Ser	Pro	Asp	Ala 25	Gly	Pro	His	Gly	Gln 30	Gly	Arg
	Val	His	Gln 35	Ala	Ala	Pro	Leu	Ser 40	Asp	Ala	Pro	His	Asp 45	Asp	Ala	His
30	Gly	Asn 50	Phe	Gln	Tyr	Asp	His 55	Glu	Ala	Phe	Leu	Gly 60	Arg	Glu	Val	Ala
35	Lys 65	Glu	Phe	Asp	Gln	Leu 70	Thr	Pro	Glu	Glu	Ser 75	Gln	Ala	Arg	Leu	Gly 80
33	Arg	Ile	Val	Asp	Arg 85	Met	Asp	Arg	Ala	Gly 90	Asp	Gly	Asp	Gly	Trp 95	Val
40	Ser	Leu	Ala	Glu 100	Leu	Arg	Ala	Trp	Ile 105	Ala	His	Thr	Gln	Gln 110	Arg	His
	Ile	Arg	Asp 115	Ser	Val	Ser	Ala	Ala 120	Trp	Asp	Thr	Tyr	Asp 125	Thr	Asp	Arg
45	Asp	Gly 130	Arg	Val	Gly	Trp	Glu 135	Glu	Leu	Arg	Asn	Ala 140	Thr	Tyr	Gly	His
50	Тут 145	Ala	Pro	Gly	Glu	Glu 150	Phe	His	Asp	Val	Glu 155	Asp	Ala	Glu	Thr	Туг 160
30	Lys	Lys	Met	Leu	Ala 165	Arg	Asp	Glu	Arg	Arg 170	Phe	Arg	Val	Ala	Asp 175	Gln
55	Asp	Gly	Asp	Ser 180	Met	Ala	Thr	Arg	Glu 185	Glu	Leu	Thr	Ala	Phe 190	Leu	His
	Pro	Glu	Glu 195	Phe	Pro	His	Met	Arg 200	Asp	Ile	Val	Ile	Ala 205	Glu	Thr	Leu
60	Glu	Asp	Leu	Asp	Arg	Asn	Lys	Asp	Gly	Tyr	Val	Gln	Val	Glu	Glu	Tvr

		210					215					220				
5	Ile 225	Ala	Asp	Leu	Tyr	Ser 230	Ala	Glu	Pro	Gly	Glu 235	Glu	Glu	Pro	Ala	Trp 240
,	Val	Gln	Thr	Glu	Arg 245	Gln	Gln	Phe	Arg	Asp 250	Phe	Arg	Asp	Leu •	Asn 255	Lys
10	Asp	Gly	His	Leu 260	Asp	Gly	Ser	Glu	Val 265	Gly	His	Trp	Val	Leu 270	Pro	Pro
	Ala	Gln	Asp 275	Gln	Pro	Leu	Val	Glu 280	Ala	Asn	His	Leu	Leu 285	His	Glu	Ser
15	Asp	Thr 290	Asp	Lys	Asp	Gly	Arg 295	Leu	Ser	Lys	Ala	Xaa 300	Ile	Leu	Gly	Asn
20	Trp 305	Asn	Met	Phe	Val	Gly 310	Ser	Gln	Ala	Thr	Asn 315	Tyr	Gly	Glu	Asp	Leu 320
	Thr	Arg	His	His	Asp 325	Glu	Leu	Xaa								
25	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO: :	171:							
30				(A) L B) T D) T	ENGT YPE : OPOL	H: 6 ami OGY:	9 am no a lin	ino cid ear	acid		: 17	1:			
35	Met 1	Cys	Trp	Leu	Arg 5	Ala	Trp	Xaa	Gln	Ile 10	Xaa	Leu	Pro	Val	Phe 15	Xaa
	Ser	Xaa	Phe	Leu 20	Ile	Gln	Leu	Leu	Ile 25	Ser	Phe	Ser	Glu	Asn 30	Gly	Phe
40	Ile	His	Ser 35	Pro	Arg	Asn	Asn	Gln 40	Lys	Pro	Arg	Asp	Gly 45	Asn	Xaa	Glu
45		50					Ser 55		Gln	Leu	Cys	Thr 60		Asp	Lys	Lys
50	(2)	INF	ORMA	TION	FOR	SEQ	D	NO:	172:							
 55				((A) I (B) 1 (D) 1	LENGT TYPE : TOPOI	TH: I am: LOGY:	l60 a ino a : lir	amino acid near	o aci): 17	72:			
60	Met 1		Leu	Phe	Ile 5		Leu	Ser	Leu	Ala		ılle	Ser	Asp	Ala 15	Met

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	Val	Met	Asp	Glu 20	Lys	Val	Lys	Arg	Ser 25	Phe	Val	Leu	Asp	Thr 30	Ala	Ser
5	Ala	Ile	Cys 35	Asn	Tyr	Asn	Ala	His 40	Tyr	Lys	Asn	His	Pro 45	Lys	Tyr	Trp
10	Cys	Arg 50	Gly	Tyr	Phe	Arg	Asp 55	Tyr	Cys	Asn	Ile	Ile 60	Ala	Phe	Ser	Pro
	Asn 65	Ser	Thr	Asn	His	Val 70	Ala	Leu	Lys	Asp	Thr 75	Gly	Asn	Gln	Leu	Ile 80
15	Val	Thr	Met	Ser	Cys 85	Leu	Asn	Lys	Glu	Asp 90	Thr	Gly	Trp	Tyr	Trp 95	Cys
	Gly	Ile	Gln	Arg 100	Asp	Phe	Ala	Arg	Asp 105	Asp	Met	Asp	Phe	Thr 110	Glu	Leu
20	Ile	Val	Thr 115	Asp	Asp	Lys	Gly	Thr 120	Trp	Pro	Met	Thr	Leu 125	Val	Trp	Glu
25	Arg	Leu 130	Ser	Gly	Thr	Lys	Pro 135	Glu	Ala	Ala	Arg	Leu 140	Pro	Lys	Leu	Ser
	Ala 145	Arg	Leu	Thr	Ala	Pro 150	Gly	Arg	Pro	Phe	Ser 155	Ser	Phe	Ala	Tyr	Xaa 160
30																
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	173:							
35			(i)	((A) I	CHA ENGI	H: 1	.23 a	mino		.ds					
40			(xi)	((D) 1	YPE: OPOL E DE	.OGY :	lir	ear	EQ I	D NC): 17	3 :			
	Met 1	Ala	. Xaa	His	Phe 5		Leu	Val	Ala	Leu 10		Ser	Val	Pro	His 15	
45	Pro	His	Leu	Leu 20		Glu	Glu	His	Lys 25		Cys	Lys	Val	Ser		Phe
50	Ser	Gly	Val		Leu	Val	Thr	Ser 40		Gln	Asp	Ser	Ser 45		Тут	Val
50	Pro	Val		Thr	: Leu	Phe	Ile 55		Leu	Gly	/ Pro	Trp 60		Trp) Asp	Leu
55	Xaa 65		Cys	Thr	Ala	Glu 70		Pro	Glu	ı Ala	a Glu 75		sei	Leu	ı Arç	Leu 80
	Cys	His	s Ser	His	Let 89		Arç	g Xaa	a Asr	va] 90		Pro	Sei	c Glr	a Ala 99	a Ala
60	Glu	Gly	/ Xaa	a Xaa	a Xaa	a Arc	Gly	/ Cys	s Glr	n His	s Arg	g Gly	/ Se:	r Arg	g Glu	ı Leu

300

100 105 110 Thr Phe Leu Ser Ala Glu Asn Glu Ala Gly Ile 115 120 5 (2) INFORMATION FOR SEQ ID NO: 174: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 129 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 174: 15 Met Lys Val Gly Ala Arg Ile Arg Val Lys Met Ser Val Asn Lys Ala 10 His Pro Val Val Ser Thr His Trp Arg Trp Pro Ala Glu Trp Pro Gln 20 25 Met Phe Leu His Leu Ala Gln Glu Pro Arg Thr Glu Val Lys Ser Arg 25 Pro Leu Gly Leu Ala Gly Phe Ile Arg Gln Asp Ser Lys Thr Arg Lys pro Leu Glu Glu Thr Ile Met Ser Ala Ala Asp Thr Ala Leu Trp 30 Pro Tyr Gly His Gly Asn Arg Glu His Gln Glu Asn Glu Leu Gln Lys Tyr Leu Gln Tyr Lys Asp Met His Leu Leu Asp Ser Gly Gln Ser Leu 35 Gly His Thr His Thr Leu Gln Gly Ser His Asn Leu Thr Ala Leu Asn 120 40 Ile 45 (2) INFORMATION FOR SEQ ID NO: 175: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 372 amino acids (B) TYPE: amino acid 50 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 175: Met Ala Tyr His Ser Phe Leu Val Glu Pro Ile Ser Cys His Ala Trp 5 10 55 . Asn Lys Asp Arg Thr Gln Ile Ala Ile Cys Pro Asn Asn His Glu Val His Ile Tyr Glu Lys Ser Gly Ala Lys Trp Thr Lys Val His Glu Leu 60 35

	цуѕ	50	ura	ASII	Gly	GIII	55	THE	GIY	ııe	Asp	60	Ala	Pro	Glu	Ser
5	Asn 65	Arg	Ile	Val	Thr	Cys 70	Gly	Thr	Asp	Arg	Asn 75	Ala	Tyr	Val	Trp	Thr 80
10	Leu	Lys	Gly	Arg	Thr 85	Trp	Lys	Pro	Thr	Leu 90	Val	Ile	Leu	Arg	Ile 95	Asn
- 0	Arg	Ala	Ala	Arg 100	Cys	Val	Arg	Trp	Ala 105	Pro	Asn	Glu	Asn	Lys 110	Phe	Ala
15	Val	Gly	Ser 115	Gly	Ser	Arg	Val	Ile 120	Ser	Ile	Cys	Tyr	Phe 125	Glu	Gln	Glu
	Asn	Asp 130	Trp	·Trp	Val	Cys	Lys 135	His	Ile	Lys	Lys	Pro 140	Ile	Arg	Ser	Thr
20	Val 145	Leu	Ser	Leu	Asp	Trp 150	His	Pro	Asn	Asn	Val 155	Leu	Leu	Ala	Ala	Gly 160
25	Ser	Cys	Asp	Phe	Lys 165	Cys	Arg	Ile	Phe	Ser 170	Ala	Tyr	Ile	Lys	Glu 175	Val
	Glu	Glu	Arg	Pro 180	Ala	Pro	Thr	Pro	Trp 185	Gly	Ser	Lys	Met	Pro 190	Phe	Gly
30	Glu	Leu	Met 195	Phe	Glu	Ser	Ser	Ser 200	Ser	Cys	Gly	Trp	Val 205	His	Gly	Val
	Cys	Phe 210	Ser	Ala	Ser	Gly	Ser 215	Arg	Val	Ala	Trp	Val 220	Ser	His	Asp	Ser
35	Thr 225	Val	Cys	Leu	Ala	Asp 230	Ala	Asp	Lys	Lys	Met 235	Ala	Val	Ala	Thr	Leu 240
40	Ala	Ser	Glu	Thr	Leu 245	Pro	Leu	Leu	Ala	Leu 250	Thr	Phe	Ile	Thr	Asp 255	Asn
	Ser	Leu	Val	Ala 260	Ala	Gly	His	Asp	Cys 265	Phe	Pro	Val	Leu	Phe 270	Thr	Tyr
45	Asp	Ala	Ala 275	Ala	Gly	Met	Leu	Ser 280		Gly	Gly	Arg	Leu 285	Asp	Val	Pro
	Lys	Gln 290	Ser	Ser	Gln	Arg	Gly 295	Leu	Thr	Ala	Arg	Glu 300	Arg	Phe	Gln	Asn
50	Leu 305	qzA	Lys	Lys	Ala	Ser 310	Ser	Glu	Gly	Gly	Thr 315	Ala	Ala	Gly	Ala	Gly 320
55	Leu	Asp	Ser	Leu	His 325	Lys	Asn	Ser	Val	Ser 330	Gln	Ile	Ser	Val	Leu 335	Ser
	Gly	Gly	Lys	Ala 340	Lys	Суѕ	Ser	Gln	Phe 345		Thr	Thr	Gly	Met 350	Asp	Gly
60	Gly	Met	Ser 355	Ile	Trp	Asp	Val	Lys 360	Ser	Leu	Glu	Ser	Ala 365	Leu	Lys	Asp

302

Leu Lys Ile Lys 370 5 (2) INFORMATION FOR SEQ ID NO: 176: (i) SEQUENCE CHARACTERISTICS: 10 (A) LENGTH: 216 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 176: Met Trp Ser Ile Gly Ala Gly Ala Leu Gly Ala Ala Ala Leu Ala Leu 15 Leu Leu Ala Asn Thr Asp Val Phe Leu Ser Lys Pro Gln Lys Ala Ala 20 Leu Glu Tyr Leu Glu Asp Ile Asp Leu Lys Thr Leu Glu Lys Glu Pro 40 Arg Thr Phe Lys Ala Lys Glu Leu Trp Glu Lys Asn Gly Ala Val Ile 25 55 Met Ala Val Arg Arg Pro Gly Cys Phe Leu Cys Arg Glu Glu Ala Ala 30 Asp Leu Ser Ser Leu Lys Ser Met Leu Asp Gln Leu Gly Val Pro Leu Tyr Ala Val Val Lys Glu His Ile Arg Thr Glu Val Lys Asp Phe Gln 35 Pro Tyr Phe Lys Gly Glu Ile Phe Leu Asp Glu Lys Lys Lys Phe Tyr Gly Pro Gln Arg Arg Lys Met Met Phe Met Gly Phe Ile Arg Leu Gly 40 Val Trp Tyr Asn Phe Phe Arg Ala Trp Asn Gly Gly Phe Ser Gly Asn 45 Leu Glu Gly Glu Gly Phe Ile Leu Gly Gly Val Phe Val Val Gly Ser 170 Gly Lys Gln Gly Ile Leu Leu Glu His Arg Glu Lys Glu Phe Gly Asp 50 Lys Val Asn Leu Leu Ser Val Leu Glu Ala Ala Lys Met Ile Lys Pro 200 Gln Thr Leu Ala Ser Glu Lys Lys 55 210

(2) INFORMATION FOR SEQ ID NO: 177:

				(E	3) TY	PE:	amir	ami no ac	id	acids	5					
5			(xi)					line		EQ II	ON O	177	:			
	Met 1	Lys	Pro	Val	Ser 5	Arg	Arg	Thr	Leu	Asp 10	Trp	Ile '	ľyr	Ser	Val 1	Leu
10	Leu	Leu	Ala	Ile 20	Val	Leu	Ile	Ser	Trp 25	Gly	Cys	Ile	Ile	Tyr . 30	Ala :	Ser
15	Met	Val	Ser 35	Ala	Arg	Arg	Gln	Leu 40	Arg	Lys	Lys	Tyr	Pro 45	Asp	Lys	Ile
	Phe	Gly 50	Thr	Asn	Glu	Asn	Leu 55									
20	(2)	INF	ORMA'	rion	FOR	SEQ	ID N	√O: 1	.78:		•					
25				(1	A) L B) T D) T	ENGT: YPE: OPOL	H: 2 ami OGY:	3 am no a lin	ino cid ear	acid		: 178	ı:			
30	Met 1	Ala	Ala	Asn	Thr 5	Phe	Val	Leu	Ile	Met 10	Gly	Ile	Pro	Thr	Ser 15	Ala
35	Asn	Ala	Xaa	Arg 20	Asp	Leu	Phe									
	(2)	INF	ORMA	TION	FOR	SEQ	ID I	NO: 3	L79 :							
40				(A) L B) T D) T	ENGT YPE : OPOL	H: 1 ami OGY:	.03 a no a lin	mino cid ear	aci		: 179	9 :			
45	Met 1					Arg						Leu		Ala	Gly 15	Val
50	Ser	Leu	. Phe	Gly 20	Met	Ser	Ala	Leu	Leu 25		Pro	Gly	Asn	Phe 30	Glu	Ser
50	Тух	Leu	Glu 35		Val	Lys	Ser	Leu 40		Leu	Gly	Pro	Ala 45		Ile	His
55	Thr	Ala 50		: Phe	Ala	Leu	Val 55		Pro	Leu	Met	туr 60	His	Thr	Trp	Asn
	Gly 65		e Arç	, His	Leu	Met 70		Asp	Leu	Gly	7 Lys	Gly	Leu	Lys	Ile	Pro 80
60	Glr	ı Lev	тут	Gln	Ser	Gly	' Val	Val	Val	. Leu	val	Leu	Thr	Val	Leu	Ser

(i) SEQUENCE CHARACTERISTICS:

					85					90					95	
5	Ser	Met	Gly	Leu 100	Ala	Ala	Met					,	,			
	(2)	INF	ORMA!	rion	FOR	SEQ	ID I	10 : 3	180:					•		
10			(i) ·	(ENGT YPE:	H: 4 ami	8 am no a	ino cid		s					
15			(xi)													
	Met 1	Thr	Lys	Ala	Ser 5	Ser	Leu	Trp	Pro	Leu 10	Lys	Thr	Thr	Cys	Gln 15	Ile
20	Ser	Gly	Thr	Val 20	Phe	Phe	Phe	Leu	Phe 25	Leu	Phe	Ser	Cys	Phe 30	Leu	Met
	Gln	Ala	Gln 35	Cys	Asp	Lys	Phe	Val 40	Gly	Trp	Asp	Phe	Phe 45	Phe	Phe	Leu
25																
30	(2)		ORMA:	SEQUI	ENCE A) L	CHAI ENGT	RACTI H: 9	ERIS' 6 am	TICS ino		s					
35			(xi)	(B) T D) T UENC	OPOL	OGY:	lin	ear	EQ II	ои с	: 18	1:			
40	Met 1	Arg	Arg	Ala	Leu 5	Ile	Pro	Pro	Cys	Arg 10	Gly	Gly	Pro	Ser	Ala 15	Ser
	Asp	Xaa	Cys	Cys 20	Ser	Cys	Ser	Pro	Ser 25	Gly	P'ne	Ser	Ala	Gly 30		Gly
45	Arg	Cys	Pro 35	Val	Gln	Gly	Cys	Leu 40	Arg	Pro	His	Arg	Val 45	Gln	Leu	Leu
	Arg	Arg 50	Trp	Gly	Pro	Gly	Ser 55	Pro	Ala	Gly	Gln	Arg 60	Leu	Ser	Lys	Gly
50	Phe 65	Gln	Leu	Leu	Arg	Trp 70	Trp	Gly	Pro	Gly	Ser 75	Pro	Ala	Pro	Glu	Pro 80
55	Arg	Lys	Gly	Pro	Phe 85	Pro	Pro	Pro	Àsp	Pro 90	Pro	Trp	Pro	Val	Thr 95	Leu

	(2)	INF	ORMAT	NOI	FOR	SEQ	ID 1	10: 1	182:							
5				C	A) L B) T D) T	ENGT: YPE: OPOL	H: 9 ami OGY:	5 am no a lin	ino cid ear	acid		: 18	. 2 :			
10	Met 1	Leu	Glu	Thr	Thr 5	Lys	His	Val	Gln	Ile 10	Ala	Cys	Met	Leu	Leu 15	Leu
	Thr	Cys	Gln	Ile 20	Phe	Leu	Pro	Ser	Ser 25	Leu	Ser	Pro	Ser	Phe 30	Ile	His
15	Ser	Leu	Thr 35	Asp	Ser	Phe	Ile	Pro 40	Leu	Lys	Lys	Leu	Тут 45	Val	Cys	Phe
20	Val	Gln 50	Ser	Thr	Leu	Leu	Lys 55	Ala	Ala	Gly	Tyr	Lys 60	Ser	Ile	Ser	Glu
	Ala 65	Leu	Gly	Phe	Asp	Xaa 70	Leu	Leu	Cys	Ser	Ser 75	Ala	Arg	Phe	Val	Trp 80
25	Ile	Cys	His	Thr	Tyr 85	Ser	Arg	Pro	Leu	Val 90	Thr	Cys	Ala	Leu	His 95	
30	(2)	INF		TION SEQU		-				:		:				
35			(xi)	(B) 1	YPE:	ami OGY:	no a	near			v 19	3.			
55	Met 1			Ile							Val			Gly	Pro 15	
40	Gly	Val	Ser	Met 20	Asp	Glu	Lys	Lys	Lys 25		Trp	•				
45	(2)	INF		TION.												
50				1	(A) I (B) 7 (D) 1	LENGT TYPE TOPOI	TH: : : am: LOGY	ll ar ino a : lir	mino acid near	acio		D: 18	34 :			
55	Met		Gly	, Gly	Leu S		Phe	e Leu	ı Lev	Leu 10		L				
	(2)	INF	FORM	MOITA	FOF	SEÇ) ID	NO:	185	:						
60			(1)	SFO	IENIC!	- CH	מפערי	יו משיו	STTC:	2.						

				(E	3) T	PE:	amir CGY:	no a	cid	acıds	5					
5			(xi)	SEQU	ENCE	DES	CRIE	OITS	1: SI	EQ II	NO:	185	:			
-	Met 1	Phe	Ala	Asp	Phe 5	Ile	Val	Val	Thr	Ala 10	Thr	Val (Gln	Arg (Cys I 15	Pro
0	Gly	Ser	Pro	Pro 20	Leu	Ser	Glu	Ile	Leu 25	Trp	Lys	Asp (Glu	Pro 30	Phe i	Ala
	.Ile	Ser	Ser 35	His	Ala	Gly	Leu	Pro 40	Trp	Leu	Ser	Ser	Trp 45	Pro .	Ala 1	Pro
15	Pro	Trp 50	Thr	Trp	Ser	Trp	Ile 55	Ser	Arg	Arg	Arg	Glu 60	His	Gly	Arg (Gly
20	Ser 65															
	(2)	INF	ORMA'	rion	FOR	SEQ	ID N	NO: .	186:							
25			(i)	(A) L B) T	ENGT YPE:		2 am no a	ino cid	: acid	s					
30	Mot	vol		SEQ	JENC:	E DE:	SCRI	PTIO	N: S			: 186		Sor	Dro	Clyr
	1				5		FIO	vai	vai	10	Cys	1111	ьеu	Ser	15	GIY
35	Ile	qzA	Ser	Ser 20	Pro	Ser										
40	(2)	INF			ENCE A) L	CHA ENGT	RACT	ERIS	TICS	: o aci	.ds					
45			(xi)				.OGY : .SCRI			SEQ I	D NO	: 18	7:			
	Met 1	_	Val	Leu	Phe 5	Val	Ala	Ile	Phe	Ala 10		Pro	Leu	Ile	Leu 15	Gly
50	Gln	Glu	тут	Glu 20	Asp	Glu	Glu	Arg	Leu 25	-	Glu	Asp	Glu	Туr 30	Tyr	Gln
55	Val	. Val	. Тут 35		Тут	Thr	.Val	Thr 40		Ser	Tyr	Asp	Asp 45	Phe	Ser	Ala
	Asp	Phe 50		: Ile	Asp	Tyr	Ser 55		Phe	e Glu	ı Ser	Glu 60	_	Arg	Leu	Asn
60	Arg 65		ı Asp	Lys	Asp	70		Glu	ı Ala	a Ile	Glu 75		Thr	Ile	Ser	Leu 80

	Glu	Thr	Ala	Arg	Ala 85	Asp	His	Pro	Lys	Pro 90	Val	Thr	Val	Lys	Pro 95	Val
5	Thr	Thr	Glu	Pro 100	Gln	Ser	Pro	Asp	Leu 105	Asn	Asp	Ala	Val	Ser 110	Ser	Leu
10	Arg	Ser	Pro 115	Ile	Pro	Leu	Leu	Leu 120	Ser	Cys	Ala	Phe	Val 125	Gln	Val	Gly
	Met	Тут 130	Phe	Met												
15	(2)	INFO	ORMA'	rion	FOR	SEQ	ID 1	NO: I	188:							
			(i)					ERIS'			c					
20			(xi)	(B) T D) T	YPE: OPOL	ami OGY:	no a lin	cid ear	•		. 18	8 :			
25			Cys		Pro					Ser				Thr		Gly
	1 Leu	Leu	Leu	Met	5 Leu	Pro	Ser	Leu	Pro	10 Ser	Pro	Ala	Ser	Gln	15 Pro	Arg
30				20					25					30		
30	PIO	Pne	Cys 35	Ser	ser	Mec	GIU	1yr 40	Phe	His	Gly	Cys	A1a 45	Ser	Pro	Ser
35	Gln	Ala 50	Ile	Ile	Gly	Gly	Phe 55	Pro	Phe	Ala	Ser	Val 60	Ala	Leu	Ala	Asp
	Ile 65	Leu	Cys	Leu	Gln											
40	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO: I	189:							
				SEQU	ENCE	СНА	RACT	ERIS	TICS	-						
45				(B) T	YPE:	ami	5 am no a lin	cid	acid	s					
				SEQ	UENC	E DE	SCRI	PTIO	N: S							
50	Met 1		Leu	Leu	Ser 5	Pro	Ala	Ile	Pro	Ala 10	Leu	Thr	Leu	Ile	Phe 15	Ile
	Leu	Met	Phe	Phe 20	Ser	Phe	Pro	Phe	Arg 25	Ala	His	Thr	Val	Val 30	Thr	Ile
55	Val	Ala	Ser 35	Gly	Phe	Leu	Gly	Leu 40	Ser	Pro	Leu	Cys	Gly 45			
60	(2)	INF	ORMA	TION	FOR	SEQ	ID:	NO:	190:							

(2) INFORMATION FOR SEQ ID NO: 190:

```
(i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 65 amino acids
                    (B) TYPE: amino acid
 5
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 190:
      Met Ala Phe Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro
                                         10
10
      Leu Gln Trp Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser
                                      25
      Tyr Gly Val Thr Arg Val Glu Ser Glu Lys Cys Asn Asn Leu Trp Leu
15
      Phe Leu Glu Thr Gly Gln Leu Pro Lys Asp Arg Ser Thr Asp Gln Arg
20
      Ser
       65
25
      (2) INFORMATION FOR SEQ ID NO: 191:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 50 amino acids
                     (B) TYPE: amino acid
30
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 191:
      Met Asn Leu Cly Met Ile Phe Ser Met Cys Gly Leu Met Leu Lys
35
      Leu Lys Trp Cys Ala Trp Val Ala Val Tyr Cys Ser Phe Ile Ser Phe
                                       25
      Ala Asn Ser Arg Ser Ser Glu Asp Thr Lys Gln Met Met Ser Ser Phe
40
                                   40
      Met Xaa
           50
45
      (2) INFORMATION FOR SEQ ID NO: 192:
              (i) SEQUENCE CHARACTERISTICS:
50
                     (A) LENGTH: 170 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 192:
55
      Met Leu Leu Asn Val Ala Leu Val Ala Leu Val Leu Leu Gly Ala Tyr
        1
                                           10
      Arg Leu Trp Val Arg Trp Gly Arg Arg Gly Leu Gly Ala Gly Ala Gly
                                       25
60
```

	Ala	Gly	Glu 35	Glu	Ser	Pro	Ala	Thr 40	Ser	Leu	Pro	Arg	Met 45	Lys	Lys	Arg
5	Asp	Phe 50	Ser	Leu	Glu	Gln	Leu 55	Arg	Gln	Туr	Asp	Gly 60	Ser	Arg	Asn	Pro
	Arg 65	Ile	Leu	Leu	Ala	Val 70	Asn	Gly	Lys	Val	Phe 75	Asp	Val	Thr	Lys	Gly 80
10	Ser	Lys	Phe	Tyr	Gly 85	Pro	Ala	Gly	Pro	Тут 90	Gly	Ile	Phe	Ala	Gly 95	Arg
15	Asp	Ala	Ser	Arg 100	Gly	Leu	Ala	Thr	Phe 105	Cys	Leu	Asp	Lys	Asp 110	Ala	Leu
	Arg	Asp	Glu 115	Tyr	Asp	Asp	Leu	Ser 120	Asp	Leu	Asn	Ala	Val 125	Gln	Met	Glu
20	Ser	Val 130	Arg	Glu	Trp	Glu	Met 135	Gln	Phe	Lys	Glu	Lys 140	Tyr	Asp	Tyr	Val
	Gly 145	Arg	Leu	Leu	Lys	Pro 150	Gly	Glu	Glu	Pro	Ser 155	Glu	Tyr	Thr	Asp	Glu 160
25	Glu	Asp	Thr	Lys	Asp 165	His	Asn	Lys	Gln	Asp 170						
30	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO: :	193:							
			(i)	(A) L	ENGT		6 am	TICS ino		s					
35			(xi)	(D) T	OPOL	OGY:	lin		EQ I	D NO	: 19	3:			
40	Met 1	Thr	Tyr	Phe	Ser 5	Gly	Leu	Leu	Val	Ile 10	Leu	Ala	Phe	Ala	Ala 15	Trp
	Val	Ala	Leu	Ala 20	Glu	Gly	Leu	Gly	Val 25	Ala	Val	Tyr	Ala	Ala 30	Ala	Val
45	Leu	Leu	Gly 35	Ala	Gly	Cys	Ala	Thr 40	Ile	Leu	Val	Thr	Ser 45	Leu	Ala	Met
	Thr	Ala 50	Asp	Leu	Ile	Gly	Pro 55	His	Thr	Asn	Ser	Gly 60		Ser	Cys	Thr
50	Ala 65	Pro														
55	. (2)	INF	ORMA'	TION	FOR	SEQ	ID	NO:	194:							
			(i)	((A) I	ENGI		2 an	TICS mino ncid		ls					
60					(D) 7											

310

(xi)	SEQUENCE	DESCRIPTION:	SEQ	ID	NO:	194:
------	----------	--------------	-----	----	-----	------

Met Ala Ala Gly Pro Ser Gly Cys Leu Val Pro Ala Phe Gly Leu Arg 10 5 Leu Leu Leu Ala Thr Val Leu Gln Ala Val Ser Ala Phe Gly Ala Glu 25 Phe Ser Ser Glu Ala Cys Arg Glu Leu Gly Phe Ser Ser Asn Leu Leu 10 40 Cys Ser Ser Cys Asp Leu Leu Gly Gln Phe Asn Leu Leu Gln Leu Asp 15 Pro Asp Cys Arg Gly Cys Cys Gln Glu Glu Ala Gln Phe Glu Thr Lys Lys Leu Tyr Ala Gly Ala Ile Leu Glu Val Cys Gly 20 (2) INFORMATION FOR SEQ ID NO: 195: 25 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 176 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 195: 30 Met Arg Gly Ser His Leu Arg Leu Leu Pro Tyr Leu Val Ala Ala Asn Pro Val Asn Tyr Gly Arg Pro Tyr Arg Leu Ser Cys Val Glu Ala Phe 35 Ala Ala Thr Phe Cys Ile Val Gly Phe Pro Asp Leu Ala Val Ile Leu 40 Leu Arg Lys Phe Lys Trp Gly Lys Gly Phe Leu Asp Leu Asn Arg Gln Leu Leu Asp Lys Tyr Ala Ala Cys Gly Ser Pro Glu Glu Val Leu Gln 75 45 Ala Glu Gln Glu Phe Leu Ala Asn Ala Lys Glu Ser Pro Gln Glu Glu 90 Glu Ile Asp Pro Phe Asp Val Asp Ser Gly Arg Glu Phe Gly Asn Pro 50 105 Asn Arg Pro Val Ala Ser Thr Arg Leu Pro Ser Asp Thr Asp Asp Ser 120 55 Asp Ala Ser Glu Asp Pro Gly Pro Xaa Ala Glu Arg Gly Gly Ala Ser 135 140 Ser Ser Cys Cys Glu Glu Glu Gln Thr Gln Gly Arg Gly Ala Glu Ala 155 60

Arg Ala Pro Ala Glu Val Trp Lys Gly Ile Lys Lys Arg Gln Arg Asp 165 170 175

5

(2) INFORMATION FOR SEQ ID NO: 196:

10

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 70 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear

15

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 196:

Met Ser Asn Ala Cys Lys Glu Leu Ala Ile Phe Leu Thr Thr Gly Ile
1 5 10 15

20 Val Val Ser Ala Phe Gly Leu Pro Ile Val Phe Ala Arg Ala His Leu 20 25 30

Ile Glu Trp Gly Ala Cys Ala Leu Val Leu Thr Gly Asn Thr Val Ile 35 40 45

25

Phe Ala Thr Ile Leu Gly Phe Phe Leu Val Phe Gly Ser Asn Asp Asp 50 55 60

Phe Ser Trp Gln Gln Trp 65 70

(2) INFORMATION FOR SEQ ID NO: 197:

35

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear

40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 197:

45 Asn Ser Gly Gly Ser Phe Pro Val Arg 20 25

50 (2) INFORMATION FOR SEQ ID NO: 198:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 73 amino acids
- (B) TYPE: amino acid
- 55 (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 198:

Met Lys Gly Glu Leu Leu Pro Phe Leu Phe Leu Thr Val Trp Leu Trp 1 5 10 15

	Leu	Tyr	Lys	Leu 20	Xaa	Phe	Gly	Glu	Ser 25	Pro	Arg	Tyr	Pro	Asn 30	Val	Ile
5	Gly	Lys	Thr 35	Tyr	Phe	Phe	Phe	Trp 40	Thr	Asp	Gln	Ile	Ser 45	Arg	Glu	Ser
	Arg	Phe 50	Leu	Glu	Arg	Leu	Ala 55	Phe	Ile	Val	Ser	Glu 60	Asn	Cye	Leu	Ile
10	Phe 65	Leu	Ile	His	Ala	Ile 70	Thr	Gly	Gln						·	
15	(2)	INF	ORMAT													
20			(i) : (xi)	()	A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami OGY:	89 a no a lin	mino cid ear	aci		: 19	9:			
25	Met 1	Ser	Gly	Phe	Ser S	Thr	Glu	Glu	Arg	Ala 10	Ala	Pro	Phe	Ser	Leu 15	Glu
	Tyr	Arg	Val	Phe 20	Leu	Lys	Asn	Glu	Lys 25	Gly	Gln	Tyr	Ile	Ser 30	Pro	Phe
30	His	Asp	Ile 35	Pro	Ile	Tyr	Ala	Asp 40	Lys	Asp	Val	Phe	His 45	Met	Val	Val
	Glu	Val 50	Pro	Arg	Trp	Ser	Asn 55	Ala	Lys	Met	Glu	Ile 60	Ala	Thr	Lys	Asp
35	Pro 65		Asn	Pro	Ile	Lys 70		Asp	Val	Lys	Lys 75	Gly	Lys	Leu	Arg	Tyr 80
40	Val	Ala	Asn	Leu	Phe 85	Pro	Tyr	Lys	Gly	Tyr 90	Ile	Trp	Asn	Tyr	Gly 95	Ala
	Ile	Pro	Gln	Thr 100		Glu	Asp	Pro	Gly 105		Asn	Asp	Lys	His 110		Gly
45	Cys	Cys	Gly 115		Asn	Asp	Pro	Ile 120	-	Val	Cys	Glu	11e 125		Ser	Lys
	Val	. Cys	: Ala	. Arg	Gly	Glu	Ile 135		Gly	Val	Lys	Val		Gly	·Ile	Leu
50	Ala 145		: Ile	Asp	Glu	Gly 150		Thr	Asp	Trp	Lys 155		Il∈	Ala	ılle	Asn 160
55	Val	. Asp	Asp	Pro	Asp 165		a Ala	. Asn	тут	170	-	Il∈	e Asr	a Asp	Va]	_
JJ	Arg	j Lev	ı Lys	Pro 180		туг	: Leu	ı Glu	Ala 185		Val	. Asp	o Trp	Phe 190		, Arg
60	Тут	. Lys	Val) Asp	Gly	/ Lys	200		ı Asr	ı Glı	ı Phe	e Ala 209		e Asr	n Ala

	0.1 0	210	2,2				215			nop		220	2,0			
5	Asp 225	His	Trp	Lys	Ala	Leu 230	Val	Thr	Lys	Lys	Thr 235	Asn	Gly	Lys	Gly	Ile 240
10	Ser	Cys	Met	Asn	Thr 245	Thr	Leu	Ser	Glu	Ser 250	Pro	Phe	Lys	Cys	Asp 255	Pro
10	Asp	Ala	Ala	, Arg 260	Ala	Ile	Val	Asp	Ala 265	Leu	Pro	Pro	Pro	Cys 270	Glu	Ser
15	Ala	Cys	Thr 275	Val	Pro	Thr	Asp	Val 280	Asp	Lys	Trp	Phe	His 285	His	Gln	Lys
	Asn															
20																
	(2)	INF	TAMSC	rion	FOR	SEQ	ID 1	VO : 2	200:							
25 .			(i) s	(A) L B) T D) T	ENGT YPE : OPOL	H: 6 ami OGY:	25 a no a lin	mino cid ear	aci		: 20	0 :			
30	Met 1	Glu	Ile	Pro	Gly 5	Ser	Leu	Cys	Lys	Lys 10	Val	Lys	Leu	Ser	Asn 15	Asn
35	Ala	Gln	Asn	Trp 20	Gly	Met	Gln	Arg	Ala 25	Thr	Asn	Val	Thr	Туг 30	Gln	Ala
33	His	His	Val 35	Ser	Arg	Asn	Lys	Arg 40	Gly	Gln	Val	Val	Gly 45	Thr	Arg	Gly
40	Gly	Phe 50	Arg	Gly	Cys	Thr	Val 55	Ттр	Leu	Thr	Gly	Leu 60		Gly	Ala	Gly
	Lys 65		Thr	Val	Ser	Met 70	Ala	Leu	Glu	Glu	Tyr 75	Leu	Val	Cys	His	Gly 80
45	Ile	Pro	Cys	Tyr	Thr 85		Asp	Gly	Asp	Asn 90	Ile	Arg	Gln	Gly	Leu 95	
50	Lys	Asn	Leu	Gly 100		Ser	Pro	Glu	Asp 105	_	Glu	Glu	Asn	Val	-	Arg
	Ile	· Ala	Glu 115		Ala	Lys	Leu	Phe 120		Asp	Ala	Gly	Leu 125		. Cys	Ile
55	Thr	Ser 130	Phe	Ile	e Ser	Pro	Tyr 135		Gln	Asp	Arg	Asr 140		Alā	Arg	Gln
	Ile 145		: Glu	Gly	⁄ Ala	Ser 150		Pro	Phe	Phe	Glu 155		. Phe	· Val	Asp	Ala 160
60	Pro	Le.	ı His	Val	. Cys	Glu	Gln	Arg	Asp	Val	Lvs	Gly	/ Leu	тул	Lys	Lys

					165					170					175	
5	Ala	Arg	Ala	Gly 180	Glu	Ile	Lys	Gly	Phe 185	Thr	Gly	Ile	Asp	Ser 190	Glu	Tyr
J	Glu	Lys	Pro 195	Glu	Ala	Pro	Glu	Leu 200	Val	Leu	Lys	Thr	Asp 205	Ser	Cys	Asp
10	Val	Asn 210	Asp	Cys	Val	Gln	Gln 215	Val	Val	Glu	Leu	Leu 220	Gln	Glu	Arg	Asp
	Ile 225	Val	Pro	Val	Asp	Ala 230	Ser	Tyr	Glu	Val	Lys 235	Glu	Leu	Tyr	Val	Pro 240
15	Glu	Asn	Lys	Leu	His 245	Leu	Ala	Lys	Thr	Asp 250	Ala	Glu	Thr	Leu	Pro 255	Ala
20	Leu	Lys	Ile	Asn 260	Lys	Val	Asp	Met	Gln 265	Trp	Val	Gln	Val	Leu 270	Ala	Glu
20	Gly	Trp	Ala 275	Thr	Pro	Leu	Asn	Gly 280	Phe	Met	Arg	Glu	Arg 285	Glu	Tyr	Leu
25	Gln	Cys 290	Leu	His	Phe	Asp	Суs 295	Leu	Leu	Asp	Gly	Gly 300	Val	Ile	Asn	Leu
	Ser 305	Val	Pro	Ile	Val	Leu 310	Thr	Ala	Thr	His	Glu 315	Asp	Lys	Glu	Arg	Leu 320
30	Asp	Gly	Cys	Thr	Ala 325	Phe	Ala	Leu	Met	Tyr 330	Glu	Gly	Arg	Arg	Val 335	Ala
35	Ile	Leu	Arg	Asn 340	Pro	Glu	Phe	Phe	Glu 345	His	Arg	Lys	Glu	Glu 350	Arg	Cys
	Ala	Arg	Gln 355	Trp	Gly	Thr	Thr	Cys 360	Lys	Asn	His	Pro	Tyr 365	Ile	Lys	Met
40	Val	Met 370	Glu	Gln	Gly	Asp	Trp 375	Leu	Ile	Gly	Gly	Asp 380	Leu	Gln	Val	Leu
	Asp 385		Val	Tyr	Trp	Asn 390		Gly	Leu	Asp	Gln 395		Arg	Leu	Thr	Pro 400
45	Thr	Glu	Leu	Lys	Gln 405		Phe	Lys	Asp	Met 410	Asn	Ala	Asp	Ala	Val 415	Phe
50	Ala	Phe	Gln	Leu 420		Asn	Pro	Val	His 425		Gly	His	Ala	Leu 430		Met
	Gln	Asp	Thr 435		Lys	Gln	Leu	Leu 440		Arg	Gly	Tyr	Arg 445		Pro	Val
55	Leu	Leu 450		His	Pro	Leu	Gly 455		Trp	Thr	Lys	Asp 460		Asp	Val	Pro
	Leu 465		Trp	Arg	Met	Lys 470		His	Ala	Ala	Val 475		Glu	Glu	Gly	Val 480
60	Leu	Asn	Pro	Glu	Thr	Thr	Val	. Val	Ala	Ile	e Phe	Pro	Ser	Pro	Met	. Met

					485					490					495	
5	Tyr	Ala	Gly	Pro 500	Thr	Glu	Val	Gln	Trp 505	His	Cys	Arg	Ala	Arg 510	Met	Val
	Ala	Gly	Ala 515	Asn	Phe	Tyr	Ile	Val 520	Gly	Arg	Asp	Pro	Ala 525	Gly	Met	Pro
10	His	Pro 530	Glu	Thr	Gly	Lys	Asp 535	Leu	Tyr	Glu	Pro	Ser 540	His	Gly	Ala	Lys
	Val 545	Leu	Thr	Met	Ala	Pro 550	Gly	Leu	Ile	Thr	Leu 555	Glu	Ile	Val	Pro	Phe 560
15	Arg	Val	Ala	Ala	Туг 565	Asn	Lys	Lys	Lys	Lys 570	Arg	Met	Asp	Tyr	Tyr 575	Asp
20	Ser	Glu	His	His 580	Glu	Asp	Phe	Glu	Phe 585	Ile	Ser	Gly	Thr	Arg 590	Met	Arg
	Lys	Leu	Ala 595	Arg	Glu	Gly	Gln	Lys .600	Pro	Pro	Glu	Gly	Phe 605	Met	Ala	Pro
25	Lys	Ala 610	Trp	Thr	Val	Leu	Thr 615	Glu	Tyr	Tyr	Lys	Ser 620	Leu	Glu	Lys	Ala
	Xaa 625															
30	(2)	TNE	ראשמי	noi	FOR	SEO.	TD N	1 0	0.1							
35	(2)			SEQUI () ()	ENCE A) L B) T	CHAI ENGT: YPE:	RACTI H: 6 ami		rics mino cid		ds					
40				SEQ	JENC!	E DES	SCRI	PTIO	N: SI							
40	1				5					10					Lys 15	
45	Ala	Phe	Gly	Gln 20	Lys	Pro	Pro	Leu	Ser 25	Thr	Glu	Asn	Ser	His 30	Glu	qzA
	Glu	Ser	Pro 35	Met	Lys	Asņ	Val	Ser 40	Ser	Ser	Lys	Gly	Ser 45	Pro	Ala	Pro
50	Leu	Gly 50	Val	Arg	Ser	Lys	Ser 55	Gly	Pro	Leu	Lys	Pro 60	Ala	Arg	Glu	Asp
	Ser 65	Glu	Asn	Lys	Asp	His 70	Ala	Gly	Glu	Ile	Ser 75	Ser	Leu	Pro	Phe	Pro 80
55	Gly	Val	Val	Leu	Lys 85	Pro	Ala	Ala	Ser	Arg 90	Gly	Gly	Pro	Gly	Leu 95	Ser
60	Lys	Asn	Gly	Glu 100	Glu	Lys	Lys	Glu	Asp 105	Arg	Lys	Ile	Asp	Ala 110	Ala	Lys

	Asn	Thr	Phe 115	Gln	Ser	Lys	Ile	Asn 120	Gln	Glu	Glu	Leu	Ala 125	Ser	Gly	Thr
5	Pro	Pro 130	Ala	Arg	Phe	Pro	Lys 135	Ala	Pro	Ser	Lys	Leu 140	Thr	Val	Gly	Gly
	Pro 145	Trp	Gly	Gln	Ser	Gln 150	Glu	Lys	Glu	Lys	Gly 155	qaA	Lys	Asn	Ser	Ala 160
10	Thr	Pro	Lys	Gln	Lys 165	Pro	Leu	Pro	Pro	Leu 170	Phe	Thr	Leu	Gly	Pro 175	Pro
15	Pro	Pro	Lys	Pro 180	Asn	Arg	Pro	Pro	Asn 185	Val	Asp	Leu	Thr	Lys 190	Phe	His
	Lys	Thr	Ser 195	Ser	Gly	Asn	Ser	Thr 200	Ser	Lys	Gly	Gìn	Thr 205	Ser	Tyr	Ser
20	Thr	Thr 210	Ser	Leu	Pro	Pro	Pro 215	Pro	Pro	Ser	His	Pro 220	Ala	Ser	Gln	Pro
	Pro 225	Leu	Pro	Ala	Ser	His 230	Pro	Ser	Gln	Pro	Pro 235	Val	Pro	Ser	Leu	Pro 240
25	Pro	Arg	Asn	Ile	Lys 245	Pro	Pro	Phe	Asp	Leu 250	Lys	Ser	Pro	Val	Asn 255	Glu
30	Asp	Asn	Gln	Asp 260	Gly	Val	Thr	His	Ser 265	Asp	Gly	Ala	Gly	Asn 270	Leu	Asp
	Glu	Glu	Gln 275	Asp	Ser	Glu	Gly	Glu 280	Thr	Tyr	Glu	qzA	Ile 285	Glu	Ala	Ser
35	Lys	Glu 290	Arg	Glu	Lys	Lys	Arg 295	Glu	Lys	Glu	Glu	Lys 300	Lys	Arg	Leu	Glu
	Leu 305	Glu	Lys	Lys*	Glu	Gln 310	Lys	Glu	Lys	Glu	Lys 315	Lys	Glu	Gln	Glu	Ile 320
40	Lys	Lys	Lys	Phe	Lys 325	Leu	Thr	Gly	Pro	Ile 330		Val	Ile	His	Leu 335	Ala
45	Lys	Ala	Суѕ	Cys 340	Asp	Val	Lys	Gly	Gly 345		Asn	Glu	Leu	Ser 350	Phe	Lys
	Gln	Gly	Glu 355	Gln	Ile	Glu	Ile	1le 360	Arg	Ile	Thr	qzA	Asn 365	Pro	Glu	Gly
50	Lys	Trp 370		Gly	Arg	Thr	Ala 375		Gly	Ser	Тут	Gly 380		Ile	Lys	Thr
	Thr 385		Val	Glu	Ile	Asp 390		Asp	Ser	Leu	1 Lys 395		Lys	Lys	Asp	Ser 400
55	Leu	Gly	Ala	Pro	Ser 405		Pro	Ile	Glu	Asp 410		Glr	Glu	Val	Tyr 415	
60	Asp	Val	Ala	Glu 420		Asp	Asp) Ile	Ser 425		His	: Ser	Glr.	Ser 430		' Ser

	Gly	Gly	11e 435	Phe	Pro	Pro	Pro	Pro 440	Asp	Asp	Asp	Ile	Тут 445	Asp	Gly	Ile
5	Glu	Glu 450	Glu	Asp	Ala	Asp	Asp 455	Gly	Ser	Thr	Leu	Gln 460	Val	Gln	Glu	Lys
	Ser 465	Asn	Thr	Trp	Ser	Trp 470	Gly	Ile	Leu	Lys	Met 475	Leu	Lys	Glý	Lys	Asp 480
10	Asp	Arg	Lys	Lys	Ser 485	Ile	Arg	Glu	Lys	Pro 490	Lys	Val	Ser	Asp	Ser 495	Asp
15	Asn	Asn	Glu	Gly 500	Ser	Ser	Phe	Pro	Ala 505	Pro	Pro	Lys	Gln	Leu 510	Asp	Met
	Gly	Asp	Glu 515	Val	Tyr	Asp	Asp	Val 520	Asp	Thr	Ser	Asp	Phe 525	Pro	Val	Ser
20	Ser	Ala 530	Glu	Met	Ser	Gln	Gly 535	Thr	Asn	Val	Gly	Lys 540	Ala	Lys	Thr	Glu
	Glu 545	Lys	Asp	Leu	Lys	Lys 550	Leu	Lys	Lys	Gln	Xaa 555	Lys	Xaa	Xaa	Lys	Asp 560
25	Phe	Arg	Lys	Lys	Phe 565	Lys	Тут	Asp	Gly	Glu 570	Ile	Arg	Val	Leu	Tyr 575	Ser
30	Thr	Lys	Val	Thr 580	Thr	Ser	Ile	Thr	Ser 585	Lys	Lys	Trp	Gly	Thr 590	Arg	Asp
	Leu	Gln	Val 595	Lys	Pro	Gly	Glu	Ser 600	Leu	Glu	Val	Ile	Gln 605	Thr	Thr	Asp
35	Asp	Thr 610	Lys	Val	Leu	Cys	Arg 615	Asn	Glu	Glu	Gly	Lys 620	Tyr	Gly	Tyr	Val
	Leu 625	Arg	Ser	Tyr	Leu	Ala 630	Asp	Asn	Asp	Gly	Glu 635	Ile	Tyr	qzA	Asp	Ile 640
40	Ala	qzA	Gly	Cys	Ile 645	Tyr	Asp	Asn	Asp							
15	(2)			rion												
50				(A) L B) T D) T	ENGT' YPE : OPOL	H: 5 ami: OGY:	5 am no a lin	ino cid ear	acid		: 20	2 :			
	Met 1			Pro										Pro	Val 15	Leu
55		Туr	Leu	Trp 20		Leu	Trp	Leu	Pro 25		Phe	Ser	Trp	Ile 30		Glu
50	Leu	Lys	Ala 35	Val	Leu	Arg	Asp	Asp 40		Leu	Ile	Ser	Ala 45		Ala	Trp

	Asn	Ala 50	Glu	Phe	Gln	Thr	Cys 55									
5																
	(2)	INFO	RMAT	иоп	FOR	SEQ	ID N	Ю: 2	03:							
10				() () ()	A) Li B) T D) T	ENGT YPE : OPOLA	H: 20 amin CGY:	ERIST 67 ar no ac line PTION	mino cid ear	acio		203	i:,			
15	Met 1	Val	Lys	Val	Thr 5	Phe	Asn	Ser	Ala	Leu 10	Ala	Gln	Lys	Glu	Ala 15	Lys
20	Lys	Asp	Glu	Pro 20	Lys	Ser	Gly	Glu	Glu 25	Ala	Leu	Ile	Ile	Pro 30	Pro	Asp
- 0	Ala	Val	Ala 35	Val	Asp	Cys	Lys	Asp 40	Pro	Asp	Asp	Val	Val 45	Pro	Val	Gly
25	Gln	Arg 50	Arg	Ala	Trp	Cys	Trp 55	Cys	Met	Cys	Phe	Gly 60	Leu	Ala	Phe	Met
	Leu 65	Ala	Gly	Val	Ile	Leu 70	Gly	Gly	Ala	Tyr	Leu 75	Tyr	Lys	Туг	Phe	Ala 80
30	Leu	Gln	Pro	Asp	Asp 85	Val	Tyr	Tyr	Cys	Gly 90	Ile	Lys	Tyr	Ile	Lys 95	qaA
35	Asp	Val	Ile	Leu 100	Asn	Glu	Pro	Ser	Ala 105	Asp	Ala	Pro	Ala	Ala 110	Leu	Tyr
	Gln	Thr	Ile 115	Glu	Glu	Asn	Ile	Lys 120	Ile	Phe	Glu	Glu	Glu 125	Glu	Val	Glu
40	Phe	Ile 130	Ser	Val	Pro	Val	Pro 135	Glu	Phe	Ala	Asp	Ser 140	Asp	Pro	Ala	Asn
	11e 145	Val	His	Asp	Phe	Asn 150	Lys	Lys	Leu	Thr	Ala 155	Tyr	Leu	Asp	Leu	Asn 160
45	Leu	Asp	Lys	Cys	Туг 165	Val	Ile	Pro	Leu	Asn 170	Thr	Ser	Ile	Val	Met 175	Pro
50	Pro	Arg	Asn	Leu 180		Glu	Leu	Leu	Ile 185	Asn	Ile	Lys	Ala	Gly 190	Thr	Tyr
	Leu	Pro	Gln 195	Ser	Tyr	Leu	Ile	His 200	Glu	His	Met	Val	Ile 205	Thr	Asp	Arg
55	Île	Glu 210		Ile	Asp	His	Leu 215		Phe	Phe	Ile	Туг 220	Arg	Leu	Cys	His
	Asp 225		Glu	Thr	Tyr	Lys		Gln	Arg	Arg	Glu 235		Ile	Lys	Gly	Ile 240
60	Gln	Lys	Arg	Glu	Ala	Ser	Asn	Cys	Phe	Ala	Ile	Arg	His	Phe	Glu	Asn

	Lvs	Phe	Ala	Val	Glu	Thr	Leu	Tle	Cve	Sor	Yaa					
5	_, 0			260			504	110	265	Ser	Add					
	(2)	INF	ORMA!	rion	FOR	SEQ	ID 1	NO: 2	204:							
10			(i)	(A) L B) T	CHAI ENGT YPE:	H: 3 ami	15 a no a	mino cid		ds					
15	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 204:															
13	Met 1	Asp	Leu	Arg	Gln 5	Phe	Leu	Met	Cys	Leu 10	Ser	Leu	Cys	Thr	Ala 15	Phe
20	Ala	Leu	Ser	Lys 20	Pro	Thr	Glu	Lys	Lys 25	Asp	Arg	Val	His	His 30	Glu	Pro
	Gln	Leu	Ser 35	Asp	Lys	Val	His	Asn 40	Asp	Ala	Gln	Ser	Phe 45	Asp	Tyr	Asp
25	His	Asp 50	Ala	Phe	Leu	Gly	Ala 55	Glu	Glu	Ala	Lys	Thr 60	Phe	Asp	Gln	Leu
30	Thr 65	Pro	Glu	Glu	Ser	Lys 70	Glu	Arg	Leu	Gly	Lys 75	Ile	Val	Ser	Lys	Ile 80
	Asp	Gly	Asp	Lys	Asp 85	Gly	Phe	Val	Thr	Val 90	Asp	Glu	Leu	Lys	Asp 95	Trp
35	Ile	Lys	Phe	Ala 100	Gln	Lys	Arg	Trp	Ile 105	Tyr	Glu	Asp	Val	Glu 110	Arg	Gln
	Trp	Lys	Gly 115	His	Asp	Leu	Asn	Glu 120	Asp	Gly	Leu	Val	Ser 125	Trp	Glu	Glu
40	Tyr	Lys 130	Asn	Ala	Thr	Tyr	Gly 135	Tyr	Val	Leu	Asp	Asp 140	Pro	Asp	Pro	Asp
45	Asp 145	Gly	Phe	Asn	Tyr	Lys 150	Gln	Met	Met	Val	Arg 155	Asp	Glu	Arg	Arg	Phe 160
	Lys	Met	Ala	Asp	Lys 165	Asp	Gly	Asp	Leu	Ile 170	Ala	Thr	Lys	Glu	Glu 175	Phe
50	Thr	Ala	Phe	Leu 180	His	Pro	Glu	Glu	Tyr 185	Asp	Tyr	Met	Lys	Asp 190	Ile	Val
	Val	Gln	Glu 195	Thr	Met	Glu	Asp	Ile 200	Asp	Lys	Asn	Ala	Asp 205	Gly	Phe	Ile
55	Asp	Leu 210	Glu	Glu	Tyr	Ile	Gly 215	Asp	Met	Tyr	Ser	His 220	Asp	Gly	Asn	Thr
60	Asp 225	Glu	Pro	Glu	Trp	Val 230	Lys	Thr	Glu	Arg	Glu 235	Gln	Phe	Val	Glu	Phe 240

	Arg	Asp	Lys	Asn	Arg 245	Asp	GΙΆ	Lys	Met	Asp 250	Lys	Glu	Glu		Lys 255	Asp
5	Trp	Ile	Leu	Pro 260	Ser	Asp	Tyr	Asp	His 265	Ala	Glu	Ala	Glu	Ala 270	Arg	His
	Leu	Val	Туг 275	Glu	Ser	Asp	Gln	Asn 280	Lys	Asp	Gly	Lys	Leu 285	Thr	Ŀys	Glu
10	Glu	Ile 290	Val	Asp	Lys	Tyr	Asp 295	Leu	Phe	Val	Gly	Ser 300	Gln	Ala	Thr	Asp
15	Phe 305	Gly	Glu	Ala	Leu	Val 310	Arg	His	Asp -	Glu	Phe 315					
	(2)	INF		NOIT												
20			(i) .	(ENCE A) L B) T D) T	ENGT YPE :	H: 2 ami	07 a no a	mino cid		ds					
25				SEQ						-						
	Met 1	Phe	Asp	Ala	Val 5	Leu	Ile	Leu	Leu	Leu 10	Ile	Pro	Leu	Lys	Asp 15	Lys
30	Leu	Val	Asp	Pro 20	Ile	Leu	Arg	Arg	His 25	Gly	Leu	Leu	Pro	Ser 30	Ser	Leu
	Lys	Arg	Ile 35	Ala	Val	Gly	Met	Phe 40	Phe	Val	Met	Cys	Ser 45	Ala	Phe	Ala
35	Ala	Gly 50		Leu	Glu	Ser	Lys 55	Arg	Leu	Asn	Leu	Val 60	Lys	Glu	Lys	Thr
40	Ile 65	Asn	Gln	Thr	Ile	Gly 70	Asn	Val	Val	Tyr	His 75		Ala	Asp	Leu	Ser 80
	Leu	Trp	Trp	Gln	Val 85	Pro	Gln	Tyr	Leu	Leu 90		Gly	Ile	Ser	Glu 95	Ile
45	Phe	Ala	. Ser	Ile 100		Gly	Leu	Glu	Phe 105		Tyr	Ser	Ala	Ala 110		Lys
	Ser	Met	Gln 115		Ala	Ile	Met	Gly 120		Phe	Phe	Phe	Phe 125		Gly	Val
50	Gly	Ser 130		· Val	Gly	Ser	Gly 135		. Leu	Ala	Leu	140		: Ile	Lys	Ala
55	Ile 145		Trp	Met	Ser	Ser 150		Thr	: Asp	Ph∈	Gly 155		ı Ile	Asn	Gly	Cys 160
_	Tyr	Leu	ı Asr	Tyr	туг 165		Ph∈	e Leu	ı Lev	170		a Ile	∋ Glr	ı Gly	Ala 175	Thr
60	Leu	Let	ı Leu	Ph∈		Ile	: Ile	e Ser	Val		з Тул	: Asp	, His	His 190		J Asp

	His	Gln	Arg 195	Ser	Arg	Ala	Asn	Gly 200	Val	Pro	Thr	Ser	Arg 205	Arg	Ala	
5																
	(2)	INFO	ORMA	поі	FOR	SEQ	ID N	10: 2	206:							
10			(i) :	-					rics mino		,					
10				{	B) T	YPE:	ami	no a	cid	acı	as					
			(xi)					lin PTIO	ear N: S1	EQ II	D NO	: 20	6:			
15	Met 1	Arg	Ser	Arg	Ile 5	Arg	Glu	Phe	Asp	Ser 10	Ser	Thr	Leu	Asn	Glu 15	Ser
20	Val	Arg	Asn	Thr 20	Ile	Met	Arg	Asp	Leu 25	Lys	Ala	Val	Gly	Lys 30	Lys	Phe
-	Met	His	Val 35	Leu	Tyr	Pro	Arg	Lys 40	Ser	Asn	Thr	Leu	Leu 45	Arg	Asp	Тrp
25	Asp	Leu 50	Trp	Gly	Pro	Leu	Ile 55	Leu	Cys	Val	Thr	Leu 60	Ala	Leu	Met	Leu
	Gln 65	Arg	Asp	Ser	Ala	Asp 70	Ser	Glu	Lys	Asp	Gly 75	Gly	Pro	Gln	Phe	Ala 80
30	Glu	Val	Phe	Val	Ile 85	Val	Trp	Phe	Gly	Ala 90	Val	Thr	Ile	Thr	Leu 95	Asn
35	Ser	Lys	Leu	Leu 100	Gly	Gly	Asn	Ile	Ser 105	Phe	Phe	Gln	Ser	Leu 110	Cys	Val
	Leu	Gly	Туг 115	Cys	Ile	Leu	Pro	Leu 120	Thr	Val	Ala	Met	Leu 125	Ile	Cys	Arg
40	Leu	Val 130	Leu	Leu	Ala	Asp	Pro 135	Gly	Pro	Val	Asn	Phe 140	Met	Val	Arg	Leu
	Phe 145	Val	Val	Ile	Val	Met 150	Phe	Ala	Trp	Ser	Ile 155	Val	Ala	Ser	Thr	Ala 160
45	Phe	Leu	Ala	Asp	Ser 165	Gln	Pro	Pro	Asn	Arg 170	Arg	Ala	Leu	Ala	Val 175	Tyr
50	Pro	Val	Phe	Leu 180	Phe	Tyr	Phe	Val	Ile 185	Ser	Trp	Met	Ile	Leu 190	Thr	Phe
	Thr	Pro	Gln 195	Xaa												
55	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	207 :							
			(i)	SEQU		CHA			TICS	: 						

(A) LENGTH: 331 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 207:

5	Met 1	Ala	Lys	Asp	Gln 5	Ala	Val	Glu	Asn	Ile 10	Leu	Val	Ser	Pro	Val 15	Val
	Val	Ala	Ser	Ser 20	Leu	Gly	Leu	Val	Ser 25	Leu	Gly	Gly	Lys	Ala 30	Thr	Thr
10	Ala	Ser	Gln 35	Ala	Lys	Ala	Val	Leu 40	Ser	Ala	Glu	Gln	Leu 45	Arg	Asp	Glu
15	Glu	Val 50	His	Ala	Gly	Leu	Gly 55	Glu	Leu	Leu	Arg	Ser 60	Leu	Ser	Asn	Ser
13	Thr 65	Ala	Arg	Asn	Val	Thr 70	Trp	Lys	Leu	Gly	Ser 75	Arg	Leu	Туг	Gly	Pro 80
20	Ser	Ser	Val	Ser	Phe 85	Ala	Asp	Asp	Phe	Val 90	Arg	Ser	Ser	Lys	Gln 95	His
	Tyr	Asn	Cys	Glu 100	His	Ser	Lys	Ile	Asn 105	Phe	Arg	Asp	Lys	Arg 110	Ser	Ala
25	Leu	Gln	Ser 115	Ile	Asn	Glu	Trp	Ala 120	Ala	Gln	Thr	Thr	Asp 125	Gly	Lys	Leu
30	Pro	Glu 130	Val	Thr	Lys	Asp	Val 135	Glu	Arg	Thr	Asp	Gly 140	Ala	Leu	Leu	Val
50	Asn 145	Ala	Met	Phe	Phe	Lys 150	Pro	His	Trp	Asp	Glu 155	Lys	Phe	His	His	Lys 160
35	Met	Val	Asp	Asn	Arg 165	Gly	Phe	Met	Val	Thr 170	Arg	Ser	Tyr	Thr	Val 175	Gly
	Val	Met	Met	Met 180	His	Arg	Thr	Gly	Leu 185	Туr	Asn	Tyr	Tyr	Asp 190	Asp	Glu
40	Lys	Glu	Lys 195	Leu	Gln	Ile	Val	Glu 200	Met	Pro	Leu	Ala	His 205	Lys	Leu	Ser
45	Ser	Leu 210	Ile	Ile	Leu	Met	Pro 215		His	Val	Glu	Pro 220	Leu	Glu	Arg	Leu
43	Glu 225		Leu	Leu	Thr	Lys 230		Gln	Leu	Lys	Ile 235		Met	Gly	Lys	Met 240
50	Gln	Lys	Lys	Ala	Val 245		Ile	Ser	Leu	Pro 250		Gly	Val	Val	Glu 255	Val
	Thr	His	Asp	Leu 260		Lys	His	Leu	Ala 265	-	Leu	Gly	Leu	Thr 270		Ala
55	Ile	Asp	Lys 275		Lys	Ala	. Asp	Leu 280		Arg	Met	: Ser	Gly 285		Lys	Asp
60	Leu	Туг 290	Leu	Ala	Ser	Val	Phe 295		Ala	Thr	Ala	Phe 300		Leu	. Asp	Thr

	305	Gly	na.ii	110		310	ALG	116	LILE	GIÀ	315	GIY	Vai	Arg		320
5	Val	Phe	Tyr	Ala	Asp 325	His	Pro	Phe	Ile	Ser 330	Xaa					
10	(2)			TION SEQUE	ENCE	CHAF	RACTI		ICS:		=					
15			(xi)	()	B) T D) T	YPE: OPOL	amin OGY:	no ao line	cid ear			: 208	B:		٠	
	Met 1	Cys	Met	Gln	Leu 5	Phe	Gly	Phe	Leu	Ala 10	Phe	Met	Ile	Phe	Met 15	Cys
20	Trp	Val	Gly	Asp 20	Val	Tyr	Pro	Val	Туг 25	Gln	Pro	Val	Gly	Pro 30	Lys	Gln
25	Tyr	Pro	Туг 35	Asn	Asn	Leu	Туг	Leu 40	Glu	Arg	Gly	Gly	Asp 45	Pro	Ser	Lys
	Glu	Pro 50	Glu	Arg	Val	Val	His 55	Tyr	Glu	Ile						
30	(2)	INF	ORMA'	rion	FOR	SEQ	ID i	NO: 2	209 :							
35				(A) L B) T D) T	ENGT YPE: OPOL	H: 3 ami OGY:	92 a no a lin	mino cid ear	aci		: 20	9 :			
40	Met 1	Asp	Ala	Leu	Val 5	Glu	Asp	Asp	Ile	Cys 10	Ile	Leu	Asn	His	Glu 15	Lys
	Ala	His	Lys	Arg 20	Asp	Thr	Val	Thr	Pro 25	Val	Ser	Ile	Tyr	Ser 30	Gly	Asp
45	Glu	Ser	Val 35		Ser	His	Phe	Ala 40	Leu	Val	Thr	Ala	Туг 45	Glu	Asp	Ile
50		50					55					60				Lys
	65					70					75					Glu 80
55					85					90					95	
60				Cys 100					105		Lys	Ser	Lys	Leu 110	_	Lys

			115					120					125			
5	Gln	Ser 130	Lys	Glu	Val	Glu	Leu 135	Leu	Gln	Leu	Arg	Thr 140	Glu	Val	Glu	Thr
5	Gln 145	Gln	Val	Met	Arg	Asn 150	Leu	Asn	Pro	Pro	Ser 155	Ser	Asn	Trp	Glu	Val 160
10	Glu	Lys	Leu	Ser	Cys 165	Asp	Leu	Lys	Ile	His 170	Gly	Leu	Glu	Gln	Glu 175	Leu
	Glu	Leu	Met	Arg 180	Lys	Glu	Cys	Ser	Asp 185	Leu	Lys	Ile	Glu	Leu 190	Gln	Lys
15	Ala	Lys	Gln 195	Thr	Asp	Pro	Tyr	Gln 200	Glu	Asp	Asn	Leu	Lys 205	Ser	Arg	Asp
20	Leu	Gln 210	Lys	Leu	Ser	Ile	Ser 215	Ser	Asp	Asn	Met	Gln 220	His	Ala	Tyr	Trp
20	Glu 225	Leu	Lys	Arg	Glu	Met 230	Ser	Asn	Leu	His	Leu 235	Val	Thr	Gln	Val	Gln 240
25	Ala	Glu	Leu	Leu	Arg 245	Lys	Leu	Lys	Thr	Ser 250	Thr	Ala	Ile	Lys	Lys 255	Ala
	Cys	Ala	Pro	Val 260	Gly	Cys	Ser	Glu	Asp 265	Leu	Gly	Arg	Asp	Ser 270	Thr	Lys
30	Leu	His	Leu 275	Met	Asn	Phe	Thr	Ala 280	Thr	Tyr	Thr	Arg	His 285	Pro	Pro	Leu
35	Leu	Pro 290	Asn	Gly	Lys	Ala	Leu 295	Cys	His	Thr	Thr	Ser 300	Ser	Pro	Leu	Pro
55	Gly 305		Val	Lys	Val	Leu 310	Ser	Glu	Lys	Ala	Ile 315	Leu	Gln	Ser	Trp	Thr 320
40	Asp	Asn	Glu	Arg	Ser 325		Pro	Asn	Asp	Gly 330	Thr	Cys	Phe	Gln	Glu 335	His
	Ser	Ser	Tyr	Gly 340		Asn	Ser	Leu	Glu 345		Asn	Ser	Trp	Val 350		Pro
45	Ser	Pro	Pro 355	-	Ser	Ser	Glu	Thr 360		Phe	Gly	Glu	Thr 365	_	Thr	Lys
50	Thr	Leu 370		Leu	Pro	Asn	Leu 375		Pro	Leu	His	Туг 380	Leu	ı Asp	Glr	His
50	Asn 385		ı Asn	Cys	: Leu	туг 390		Asn	ı							
55	(2)	INF	ORMA	AOIT.	I FOF	R SEÇ	DI Q	NO:	210:							

(i) SEQUENCE CHARACTERISTICS:

60

(A) LENGTH: 27 amino acids

(B) TYPE: amino acid

```
(D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 210:
      Met His His Thr Gln Leu Met Phe Ile Tyr Leu Phe Ile Tyr Leu
 5
      Phe Ile Leu Gly Val Phe Phe Phe Phe Xaa
                  20
10
      (2) INFORMATION FOR SEQ ID NO: 211:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 39 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 211:
20
     Met Asn Cys Ile Leu Leu Tyr Leu Leu Ile Pro Thr Ile Ser Ile
      Ser Val Val Pro Tyr Val Ala Leu Asn Ile Lys Tyr Ile Lys Glu Cys
25
      Thr Glu Asn Ser Phe Tyr Xaa
              35
30
      (2) INFORMATION FOR SEQ ID NO: 212:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 71 amino acids
35
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 212:
     Met Leu Leu His Leu Thr Ala Ala Phe Leu Gln Arg Ala Gln Phe Ser
40
                                          10
      Thr Tyr Phe Pro Gly Tyr Phe Asp Gly Gln Tyr Trp Leu Trp Trp Val
45
      Phe Leu Val Leu Gly Phe Leu Leu Phe Leu Arg Gly Phe Ile Asn Tyr
      Ala Lys Val Arg Lys Met Pro Glu Thr Phe Ser Asn Leu Pro Arg Thr
50
      Arg Val Leu Phe Ile Tyr Xaa
             70
55
      (2) INFORMATION FOR SEQ ID NO: 213:
             (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 83 amino acids
60
                    (B) TYPE: amino acid
```

			(xi)	SEQI		OPOLO E DES				EQ II	ONO:	213	3:			
5	Met 1													Gly	Phe 15	Phe
J		Ser	Phe	Cys 20		Thr	Thr	Ser	Ala 25		Gly	Arg	Tyr	Gly		Ile
10	Ser	Gly	Phe 35		Leu	Ser	Leu	Ile 40		Trp	Ile	Leu	Ile .45	Val	Arg	Phe
	Ser	Thr 50		Phe	Pro	Ala	Phe 55		Asn	Ser	Leu	Ser 60		Ser	Lys	Arg
15	Thr 65		Ala	Gly	Ser	Glu 70		Arg	Cys	Arg	Thr 75		Arg	Asn	Asn	His 80
20		Leu	Xaa								,3					
20																
25	(2)	INF		TION SEQU												
			(1)	(A) L B) T	ENGT YPE:	H: 8 ami	1 am	nino cid		ls					
30			(xi)	SEQ		OPOL E DE				EQ I	D NO	: 21	4 :			
	Met 1	Ser	Lys	Arg	Ser 5	Ala	Ser	Phe	Ile	Leu 10		Pro	Leu	Leu	Phe 15	Leu
35	Lys	Gly	Ser	Phe 20	Ala	Lys	Leu	Asn	Ala 25		Ile	Ser	Asp	Cys 30	Leu	Glu
40	Glu	Arg	Tyr 35		His	Asn	Leu	Trp		Val	Phe	Gln	Gly 45		Val	Ile
40	Thr	Glu 50		His	Leu	Ser	Arg 55		Ser	Lys	Thr	Leu 60		Ser	Leu	Cys
45	Tyr 65		Phe	val	Ile	Asn 70		Tyr	Ile	Phe	Phe 75		Phe	. Leu	Asp	Ile 80
	Thr															
50																
	(2)	INF	FORMA	TION	FOR	SEÇ	ID	NO:	215:							
55			(i)		(A) : (B) '	LENG	rh: 4	49 au ino	mino acid	aci	ds	•				
60	Met	: Cv) SE(_				l I.e.	ı Phe	e Ser
-	1,10,0	- ~y=		- wet					- + 1 t	~ 5.11	- CA:	116	va.	י הכנ		

	1				5					10					15	
5	Glu	Lys	Ile	Ile 20	Gln	Leu	Cys	Ala	Ser 25	Ile	Ala	Phe	Leu	Cys 30	Phe	Val
3	Lys	His	Val 35	Pro	Trp	Pro	Lys	Trp 40	Lys	Arg	Lys	Суѕ	Leu 45	Ile	Asn	Ala
10	Phe															
15	(2)	INFO	ORMA'			SEQ CHAI										
20				(A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami OGY:	03 a no a lin	mino cid ear	aci						
20			(xi)	SEQ	UENC:	E DE	SCRI:	PTIO	N: SI	EQ I	ON C	: 21	6 :			
	Met 1	Thr	Leu	Arg	Pro 5	Ser	Leu	Leu	Pro	Leu 10	His	Leu	Leu	Leu	Leu 15	Leu
25	Leu	Leu	Ser	Ala 20	Alā	Val	Cys	Arg	Ala 25	Glu	Ala	Gly	Leu	Glu 30	Thr	Glu
30	Ser	Pro	Val 35	Arg	Thr	Leu	Gln	Val 40	Glu	Thr	Leu	Val	Glu 45	Pro	Pro	Glu
50	Pro	Cys 50	Ala	Glu	Pro	Ala	Ala 55	Phe	Gly	Asp	Thr	Leu 60	His	Ile	His	Tyr
35	Thr 65	Gly	Ser	Leu	Val	Asp 70	Gly	Arg	Ile	Ile	Asp 75	Thr	Ser	Leu	Thr	Arg 80
	Asp	Pro	Leu	Val	Ile 85	Glu	Leu	Gly	Gln	Lys 90	Gln	Val	Ile	Pro	Gly 95	Leu
40	Glu	Gln	Ser	Leu 100	Leu	Asp	Met	Cys	Val 105	Gly	Glu	Lys	Arg	Arg 110	Ala	Ile
45	Ile	Pro	Ser 115	His	Leu	Ala	Тут	Gly 120	Lys	Arg	Gly	Phe	Pro 125	Pro	Ser	Val
,5	Pro	Ala 130	Asp	Ala	Val		Gln ,135	Tyr	Asp	Val	Glu	Leu 140	Ile	Ala	Leu	Ile
50	Arg 145	Ala	Asn	Tyr	Trp	Leu 150	Lys	Leu	Val	Lys	Gly 155	Ile	Leu	Pro	Leu	Val 160
	Gly	Met	Ala	Met	Val 165	Pro	Pro	Ser	Trp	Ala 170	Ser	Leu	Gly	Ile	Thr 175	Tyr
55	Thr	Glu	Arg	Pro 180	Ile	Asp	Pro	Lys	Ser 185	Pro	Lys	Arg	Ser	Ser 190	Arg	Lys
60	Arg	Asn	Glu 195	Thr	Arg	Ala	Lys	Arg 200	Asn	Asn	Lys					

	(2)	TIME	JRWAI	TOM	ron	SEQ	יו טיי	NO: 2	17:							
5			(i) S	- (. (:	A) L B) T	ENGTI YPE :	H: 1 ami		mino cid		ds					
10			(xi)	SEQ	JENCI	E DES	SCRI	PTIO	N: SI	EQ II	ON C	: 21	7:			
10	Met 1	Lys	Thr	Leu	Met 5	Thr	Ile	Cys	Pro	Gly 10	Thr	Val	Leu	Leu	Val 15	Phe
15	Ser	Ile	Ser	Leu 20	Trp	Ile	Ile	Ala	Ala 25	Trp	Thr	Val	Arg	Val 30	Cys	Glu
	Ser	Pro	Glu 35	Ser	Pro	Ala	Gln	Pro 40	Ser	Gly	Ser	Ser	Leu 45	Pro	Ala	Trp
20	Tyr	His 50	Asp	Gln	Gln	Asp	Val 55	Thr	Ser	Asn	Phe	Leu 60	Gly	Ala	Met	Trp
25	Leu 65	Ile	Ser	Ile	Thr	Phe 70	Leu	Ser	Ile	Gly	Туг 75	Gly	Asp	Met	Val	Pro 80
	His	Thr	Tyr	Cys	Gly 85	Lys	Gly	Val	Cys	Leu 90	Leu	Thr	Gly	Ile	Met 95	Gly
30	Ala	Gly	Cys	Thr 100	Ala	Leu	Val	Val	Ala 105	Val	Val	Ala	Arg	Lys 110	Leu	Glu
	Leu	Thr	Lys 115	Ala	Glu	Lys	His	Val 120	His	Xaa	Phe	Met	Met 125	Asp	Thr	Gln
35	Leu	Thr 130	Lys	Arg	Ile	Lys	Asn 135	Хаа	Ala	Ala	Asn	Val 140	Leu	Xaa	Glu	Thr
40	Trp 145	Leu	Ile	Tyr	Lys	His 150	Thr	Lys	Leu	Leu	Lys 155	Lys	Ile	Asp	His	Ala 160
	Lys	Val	Arg	Asn	Thr 165	Arg	Gly	Ser	Ser	Ser 170	-	Tyr	Pro	Pro	Val 175	Glu
45	Glu	Arg	Gln	Asp 180	Gly	Thr	Glu	Glu	Ala 185	Glu						
50	(2)	INF	ORMA						218: TICS							
- -					(B) 1	YPE:	am:	90 an ino a : lir		acio	ls					
55			(xi)	SEÇ	UENC	E DE	SCRI	PTIC	N: 5	SEQ]	D NC): 21	18:			
	Met 1		: Phe	Leu	Ala 5		Leu	Val	. Leu	Leu 10		' Val	. Ser	: Ile	Phe	. Leu
60	17-7	C		01 -		Dro	The	- mb	1	21-	. n					. D

				20					25					30		
5	Ala	Thr	Gly 35	Pro	Ala	Asp	Asp	Glu 40	Ala	Pro	Asp	Ala	Glu 45	Thr	Thr	Ala
3	Ala	Ala 50	Thr	Thr	Ala	Thr	Thr 55	Ala	Ala	Pro	Thr	Thr 60	Ala	Thr	Thr	Ala
10	Ala 65	Ser	Thr	Thr	Ala	Arg 70	Lys	Asp	Ile	Pro	Val 75	Leu	Pro	Lys	Trp	Val 80
	Gly	Asp	Leu	Pro	Asn 85	Gly	Arg	Val	Cys	Pro 90						
15																
	(2)	INF	ORMA	rion	FOR	SEQ	ID 1	NO: 2	219:							
20				(A) L B) T D) T	ENGT YPE: OPOL	H: 1 ami OGY:	39 a no a lin	mino cid ear	aci		: 21	9 :			
25	Met 1	Ser	Ser	Ala	Ala 5	Ala	qzA	His	Trp	Ala 10	Trp	Leu	Leu	Val	Leu 15	Ser
30	Phe	Val	Phe	Gly 20	Cys	Asn	Val	Leu	Arg 25	Ile	Leu	Leu	Pro	Ser 30	Phe	Ser
	Ser	Phe	Met 35	Ser	Arg	Val	Leu	Gln 40	Lys	Asp	Ala	Glu	Gln 45	Glu	Ser	Gln
35	Met	Arg 50	Ala	Glu	Ile	Gln	Asp 55	Met	Lys	Gln	Glu	Leu 60	Ser	Thr	Val	Asn
	Met 65	Met	Asp	Glu	Phe	Ala 70	Arg	Tyr	Ala	Arg	Leu 75	Glu	Arg	Lys	Ile	Asr 80
40	Lys	Met	Thr	Asp	Lys 85	Leu	Lys	Thr	His	Val 90	Lys	Ala	Arg	Thr	Ala 95	Glr
45	Leu	Ala	Lys	Ile 100	Lys	Trp	Val	Ile	Ser 105	Val	Ala	Phe	Tyr	Val 110	Leu	Glr
	Ala	Ala	Leu 115	Met	Ile	Ser	Leu	Ile 120	Trp	Lys	Tyr	Tyr	Ser 125		Pro	Val
50	Ala	Val 130		Pro	Ser	Lys	Trp 135	Ile	Thr	Leu	Xaa					
55	(2)	INF	ORMA	TION	FOR	SEQ	ID:	NO:	220:							
- •			(i)		A) [ENGT	rH: 4		nino		is					
-60			(xi)	(SEQ				lir PTIC		EO T	D NC): 27	20:			

WO 98/42738 PCT/US98/05311

	Met 1	Ser	Ser	Ala	Ala 5	Ala	Asp	His	Trp	Ala 10	Trp	Leu	Leu	Val	Leu 15	Ser
5	Phe	Val	Phe	Gly 20	Cys	Asn	Val	Leu	Arg 25	Ile	Leu	Leu	Pro	Ser 30	Phe	Ser
10	Ser	Phe	Met 35	Ser	Arg	Val	Leu	Gln 40	Lys	Asp	Ala	Asp	Arg 45	Ser	His	Arg
15	(2)	INF	ORMAT	rion	FOR	SEQ	ID I	NO: :	221:							
20				(A) L B) T D) T	ENGT YPE: OPOL	H: 7 ami OGY:	0 am no a lin	ino cid ear	acid		: 22	1:			
25	Met 1	Thr	Ala	Pro	Leu 5	Pro	Pro	Leu	Ser	Gly 10	Leu	Ala	Leu	Phe	Leu 15	Ile
	Val	Phe	Phe	Ser 20	Leu	Gly	Val	Phe	Cys 25	Ile	Cys	His	Ser	His 30	Trp	Tyr
30	His	Thr	Leu 35	Gln	Gln	Met	Ala	Gly 40		Glu	Pro	Lys	Ala 45	Leu	Leu	Leu
25	Ser	Pro 50		Ala	Ala	Thr	Thr 55	Phe	Val	Thr	Val	Thr 60		Glu	Val	Trp
35	Lys 65		Gln	Ala	Leu	Ala 70										
40	(2)	INF	'ORMA	TION	FOR	SEQ	ID	NO:	222:							
45			(i) (xi)	((A) I (B) 1 (D) 1	LENGT TYPE : TOPOI	H: 8 : am: LOGY	33 ar ino a : lin	mino acid near	acio		D: 22	22:			
50	Met		Cys	Ser	Val		Leu	Lei	ı Lev	ı Ile		ı Gly	/ Leu	ı Arç	Cys	Ser
	Gly	/ Val	L Arg	Pro 20		/ Leu	ı Val	. Gl ₃	/ Glu 25		/ His	s Asr	ı Pro	Ser 30		ı Leu
55	Val	l Cys	Leu 35		Leu	ı Lys	as,	Sei 40		g Thi	: Ası	n Glr	n Gly 45		Cys	Pro
60	Gly	/ Gly 50		Trp	Ser	r Glu	Arg 59	-	o Ile	e Glu	ı Se:	r Va:		r Sei	. Asp) Asn

	Cys 65	Glu	Ala	Thr	Leu	Gly 70	Тут	Arg	Asn	His	Ser 75	Leu	Pro	Ser	Asn	Tyr 80
5	Tyr	Asn	Ser													
10	(2)			SEQUE	ENCE	CHAF	RACTE	IO: 2 ERIST 3 ami	ICS:		5					
15			(xi)	(1	Y (C	OPOL	OGY:	no ac line PTION	ear	EQ II	O NO:	: 223	3:			
	Met 1	Leu	Thr	Arg	Ser 5	Leu	Lys	Thr	Leu	Pro 10	Ser	Ala	Cys	Thr	Ala 15	Phe
20	Leu	Leu	Leu	Phe 20	Phe	Leu	Phe	Ser	Ser 25	Gly	Asp	Pro	Glu	Leu 30	Ser	Cys
25	Ser	Cys	Thr 35	Leu	Arg	Thr	Gln	Ser 40	Ser	Trp	Ser					
	(2)	INF	ORMA	noi	FOR	SEQ	ID 1	NO: 2	24:							
30			(i)	(.	A) L	ENGT	H: 1	ERIST 84 au no a	mino		ds			•		
35				SEQ	UENC	E DE	SCRI	lin PTIO	N: S							
	Met 1		Arg	Pro	Ser 5	Val	Leu	Leu	Leu	Leu 10	Leu	Leu	Leu	Arg	His 15	Gly
40	Ala	Gln	Gly	Lys 20	Pro	Ser	Pro	Asp	Ala 25	Gly	Pro	His	Gly	Gln 30	Gly	Arg
	Val	His	Gln 35	Ala	Ala	Pro	Leu	Ser 40	Asp	Ala	Pro	His	Asp 45	Asp	Ala	His
45	Gly	Asn 50		Gln	Tyr	Asp	His 55	Glu	Ala	Phe	Leu	Gly 60	Arg	Glu	Val	Ala
50	Lys 65		Phe	Asp	Gln	Leu 70		Pro	Glu	Glu	Ser 75	Gln	Ala	Arg	Leu	Gly 80
50	Arg	Ile	val	Asp	Arg 85	Met	Asp	Arg	Ala	Gly 90	Asp	Gly	qzA	Gly	Trp 95	
55	Ser	: Lev	ı Ala	Glu 100	Leu	Arg	Ala	Trp	Ile 105		His	Thr	Gln	Gln 110		His
	Ile	e Arg	Asp 115		Val	Ser	Ala	Ala 120		Asp	Thr	Tyr	Asp 125		Asp	Arg
60	Asp	Gly	/ Arg	Val	Gly	Trp	Glu	Glu	Leu	Arg	Asn	Xaa	Thr	Tyr	Gly	His

		130					135					140				
5	Xaa 145	Xaa	Pro	Xaa	Glu	Glu 150	Phe	His	Asp	Val	Glu 155	Asp	Ala	Glu	Thr	Туг 160
3	Lys	Lys	Met	Leu	Xaa 165	Arg	Asp	Glu	Arg	Arg 170	Phe	Arg	Val	Ala	Asp 175	Gln
10	Asp	Gly	qzA	Ser 180	Met	Ala	Thr	Arg								
15	(2)	INFO		(ENCE A) L B) T	CHA ENGT YPE:	RACT. H: 7 ami	ERIS' 1 am no a	TICS ino cid		s					
20			(xi)	SEQ				lin PTIO		EQ I	D NO	: 22	5 :			
	Met 1	Trp	Leu	Phe	Ile 5	Leu	Leu	Ser	Leu	Ala 10	Leu	Ile	Ser	Asp	Ala 15	Met
25	Val	Met	Asp	Glu 20	Lys	Val	Lys	Arg	Ser 25	Leu	Cys	Trp	Thr	Arg 30	Leu	Leu
30	Pro	Ser	Ala 35	Thr	Thr	Met	Pro	Хаа 40	Thr	Arg	Ile	Thr	Pro 45	Asn	Thr	Gly
	Ala	Glu 50	Xaa	Ile	Ser	Val	Xaa 55	Thr	Ala	Thr	Ser	Ser 60	Pro	Ser	Pro	Leu
35	Thr 65	Ala	Pro	Ile	Met	Trp 70	Pro									
40	(2)	INF			ENCE A) I	CHA	RACT	ERIS	TICS uino	acid	ls					
45			(xi)		D) 1	OPOL	OGY :	no a lin PTIO	ear		D NO	ı: 22	6:			
	Met 1		Val	Phe	Val 5		Glu	Ile	Phe	Leu 10						
50																
	(2)	INF	ORMA:	TION	FOR	SEQ	ID	NO:	227 :							
55				•	(A) I (B) 3 (D) 3	LENGT TYPE : TOPOI	TH: 3 ami	l38 á ino á : lir	mind acid near	aci): 22	27 :			
60	Met	Ala												Let	Ala	Leu

	1				5					10					15	
5	Thr	Phe	Ile	Thr 20	Asp _.	Asn	Ser	Leu	Val 25	Ala	Ala	Gly	His	Asp 30	Cys	Phe
J	Pro	Val	Leu 35	Phe	Thr	Tyr	Asp	Ala 40	Ala	Ala	Gly	Met	Leu 45	Ser	Phe	Gly
10	Gly	Arg 50	Leu	Asp	Val	Pro	Lys 55	Gln	Ser	Ser	Gln	Arg 60	Gly	Leu	Thr	Ala
	Arg 65	Glu	Arg	Phe	Gln	Asn 70	Leu	Asp	Lys	Lys	Ala 75	Ser	Ser	Glu	Gly _.	Gly 80
15	Thr	Ala	Ala	Gly	Ala 85	Gly	Leu	Asp	Ser	Leu 90	His	Lys	Asn	Ser	Val 95	Ser
20	Gln	Ile	Ser	Val 100	Leu	Ser	Gly	Gly	Lys 105	Ala	Lys	Cys	Ser	Gln 110	Phe	Cys
-0	Thr	Thr	Gly 115	Met	Asp	Gly	Gly	Met 120	Ser	Ile	Trp	Asp	Val 125	Lys	Ser	Leu
25	Glu	Ser 130	Ala	Leu	Lys	Asp	Leu 135	Lys	Ile	Lys						
30	(2)	INF				SEQ										
			(1)	(A) L		H: 2	3 am	ino	acid	ls					
35			(xi)			OPOL E DE				EQ I	D NC): 22	8 :			
	Leu 1		Ser	Leu	Ser 5	Thr	Ala	Pro	Ser	Ser 10		Leu	Pro	Thr	Leu 15	Gly
40	Ala	Arg	Arg	Thr 20	Arg	Ser	Lys									
45	(2)		004	~~~	707	cro	TD		222							
40	(2)	TNF				SEQ CHA										
5 0			(1)	-	(A) I		TH: 3	133 a	mino	o aci	ids					
50			(xi)			ropoi				SEQ I	D NO	D: 22	9 :			
55	Met 1		Туг	Phe	Ser 5		Leu	Leu	Val	. Ile		ı Alş	. Phe	Ala	Ala 15	
	Val	Ala	. Leu	Ala 20		Gly	Leu	Gly	′ Val		a Val	Туг	Ala	Ala 30	Ala	Val
60	Leu	Leu	Gly 35		Gly	Cys	Ala	Thr 40		e Lei	ı Val	L Thr	Ser 45		. Ala	Met

	Thr	Ala 50	Asp	Leu	Ile	Gly	Pro 55	His	Thr	Asn	Ser	Gly 60	Ala	Phe	Val	Tyr
5	Gly 65	Ser	Met	Ser	Phe	Leu 70	Asp	Lys	Val	Ala	Asn 75	Gly	Leu	Ala	Val	Met 80
10	Ala	Ile	Gln	Ser	Leu 85	His	Pro	Cys	Pro	Ser 90	Glu	Leu	Cys	Cys	Arg 95	Ala
.0	Cys	Val	Ser	Phe 100	Tyr	His	Trp	Ala	Met 105	Val	Ala	Val	Thr	Gly 110	Gly	Val
15	Gly	Val	Ala 115	Ala	Ala	Leu	Cys	Leu 120	Cys	Ser	Leu	Leu	Leu 125	Trp	Pro	Thr
	Arg	Leu 130	Arg	Arg	Xaa											
20		•														
	(2)	INF	ORMA	rion	FOR	SEQ	ID i	NO: 2	230:							
25			(i) : (xi)	(A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami OGY:	8 am no a lin	ino cid ear	acid		. 23	ο.			
30	Gly													T3 -	τ	Met
2.0	1	Lys	. 10	1111	5	Lys	JCI	ьец	PIO	10	Mec	пр	mec	TIE	15	Met
35	Gln	Pro	Ile	Ile 20	Met	Ile	Ser	Met	Met 25	Ser	Asn	Gly				
	(2)	INF	ORMA	rion	FOR	SEQ	ID I	NO: :	231:							
40			(i).	(A) L B) T	ENGT YPE :	H: 6 ami	ERIS l am no a lin	ino cid		s					
45			(xi)	SEQ	UENC	E DE	SCRI	PTIO	N: S	EQ I	D NO	: 23	1:			
	Met 1	Gln	Gly	Lys	Phe 5	Met	Lys	Val	Gln	Val 10	Tyr	Arg	Phe	Leu	Lys 15	
50	Leu	Leu	Met	Leu 20	Leu	Cys	Met	Phe	Val 25	Asn	Arg	Gly	Met	Ser 30	Lys	qzA
	Ser	Thr	Lys 35	Lys	Pro	Gly	Gln	Glu 40	Lys	Leu	Lys	Val	Ser 45	Leu	Gly 	Ser
55	Ile	Leu 50	Asn	Met	Lys	Ser	Gln 55	Arg	Pro	Leu	Ser	Trp 60	Cys			
60	(2)	INF	ORMA'	TION	FOR	SEQ	ID	NO :	232:							

```
(i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 29 amino acids
                    (B) TYPE: amino acid
 5
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 232:
     Met Met Glu Arg Ser Met Met Ile Leu Leu Met Ala Ala Ser Met Thr
10
     Met Thr Ser Thr Gln Leu Trp Ser Phe Cys Cys Val His
                                       25
15
      (2) INFORMATION FOR SEQ ID NO: 233:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 18 amino acids
20
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 233:
     Met Trp Tyr Gln Leu Ala Lys Glu Glu Pro Gly Val Gly Ala Cys Ala
25
                                          10
     Leu Asp
30
      (2) INFORMATION FOR SEQ ID NO: 234:
             (i) SEQUENCE CHARACTERISTICS:
35
                    (A) LENGTH: 2 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 234:
40
      Leu Xaa
        1
45
      (2) INFORMATION FOR SEQ ID NO: 235:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 72 amino acids
                     (B) TYPE: amino acid
50
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 235:
      Met Leu Ile Cys Arg Leu Val Leu Leu Ala Asp Pro Gly Pro Val Asn
                        5
                                           10
55
      Phe Met Val Arg Leu Phe Val Val Ile Val Met Phe Ala Trp Ser Ile
      Val Ala Ser Thr Ala Phe Leu Ala Asp Ser Gln Pro Pro Asn Arg Arg
60
                                   40
```

	Ala	Leu 50	Ala	Val	Tyr	Pro	Va1 55	Phe	Leu	Phe	Tyr	Phe 60	Val	Ile	Ser	Trp
5	Met 65	Ile	Leu	Thr	Phe	Thr 70	Pro	Gln								
10	(2)	INF	ORMA!	rion	FOR	SEQ	ID 1	NO: 2	236:							
15				(A) L B) T D) T	ENGT YPE : OPOL	H: 9 ami OGY:	6 am no a lin	ino cid ear	acid		: 23	5 :			
20	Met 1	Arg	Ser	Leu	Leu 5	Leu	Leu	Ser	Ala	Phe 10	Cys	Leu	Leu	Glu	Ala 15	Ala
20	Leu	Ala	Ala	Glu 20	Val	Lys	Lys	Pro	Ala 25	Ala	Ala	Ala	Ala	Pro 30	Gly	Thr
25	Ala	Glu	Lys 35	Leu	Ser	Pro	Lys	Ala 40	Ala	Thr	Leu	Ala	Glu 45	Arg	Xaa	Pro
	Ala	Trp 50	Pro	Ser	Ala	Cys	Thr 55	Arg	Pro	Trp	Pro	Arg 60	Thr	Arg	Gln	Trp
30	Arg 65	Thr	Ser	Trp	Суѕ	His 70	Pro	Trp	Trp	Trp	Pro 75	Arg	Arg	Trp	Gly	Ser 80
35	Cys	Arg	Trp	Ala	Ala 85	Arg	Arg	Pro	Arg	Arg 90	Arg	Arg	Pro	Arg	Gln 95	Cys
40	(2)	INF		TION		_										
45				(A) L B) T D) T	ENGT YPE : OPOL	H: 1 ami OGY:	.43 a .no a lin	mino cid ear	aci		: 23	7:			
50	Met 1	Arg	Ser	Leu	Leu 5	Leu	Leu	Ser	Ala	Phe 10	Cys	Leu	Leu	Glu	Ala 15	Ala
	Leu	Ala	Ala	Glu 20	Val	Lys	Lys	Pro	Ala 25		Ala	Ala	Ala	Pro 30	Gly	Thr
55	Ala	Glu	Lys 35		Ser	Pro	Lys	Ala 40		Thr	Leu	Ala	Glu 45		Ļys	Arg
60	Pro	Gly 50		Gln	Leu	Val	Pro 55		His	Gly	Gln	Gly 60		Gly	Ser	Gly

	Glu 65	His	Pro	Gly	Val	Thr 70	Arg	Gly	Gly	Gly	Leu 75	Val	Ala	Gly	Ala	Arg 80
5	Val	Ala	Gly	Arg	Gln 85	Gly	Asp	His	Gly	Val 90	Ala	Gly	Gln	Gly	Ser 95	Ala
	Glu	Arg	Arg	Ala 100	Ala	Ala	Arg	Arg	Gly 105	Gly	Ala	Arg	Arg	Pro 110	Glý	Arg
10	Ala	Ala	Ala 115	Leu	Thr	Gln	Gln	Leu 120	His	Gly	Ala	Gln	Arg 125	Asp	Leu	Glu
15	Ala	Gly 130	Gln	Pro	Thr	Val	Arg 135	Thr	Gln	Leu	Ser	Glu 140	Leu	Arg	Xaa	
•	(2)	INF	ORMA!	rion	FOR	SEQ	ID I	NO: 2	238:							
20			(i) .	- (A) L B) T	ENGT YPE :	H: 1 ami	ERIS' 42 a no a lin	mino cid		ds					
25			(xi)						-	EQ I	D NO	: 23	8 :			
	Met 1	Arg	Ser	Leu	Leu 5	Leu	Leu	Ser	Ala	Phe 10	Cys	Leu	Leu	Glu	Ala 15	Ala
30	Leu	Ala	Ala	Glu 20	Val	Lys	Lys	Pro	Ala 25	Ala	Ala	Ala	Ala	Pro 30	Gly	Thr
	Ala	Glu	Lys 35	Leu	Ser	Pro	Lys	Ala 40	Ala	Thr	Leu	Ala	Glu 45	Arg	Xaa	Arg
35	Pro	Gly 50	Leu	Gln	Leu	Val	Pro 55	Gly	His	Gly	Gln	Gly 60	Pro	Gly	Ser	Gly
40	Glu 65	His	Pro	Gly	Val	Thr 70	Arg	Gly	Gly	Gly	Leu 75	Val	Ala	Gly	Ala	Arg 80
	Val	Ala	Gly	Arg	Gln 85	Gly	Asp	His	Gly	Val 90	Ala	Gly	Gln	Gly	Ser 95	Ala
45	Glu	Arg	Arg	Ala 100		Ala	Arg	Arg	Gly 105		Ala	Arg	Arg	Pro 110	Gly	Arg
	Ala	Ala	Ala 115		Thr	Gln	Gln	Leu 120		Gly	Ala	Gln	Arg 125		Leu	Glu
50	Ala	Gly 130	Gln	Pro	Thr	Val	Arg 135		Gln	. Leu	Ser	Glu 140		Arg		
55	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	239:							
			(i)	_	(A) I	LENG	PH: 5	TERIS	nino		ds					
60								ino a : lir								

			(YT)	SEQU) LIVC	5 1/15	JCICIL	. 1 101	N. DI	יו על	יטאו כ	: 43) :			
5	Asp 1	Pro	Glu	Ala	Ala 5	Asp	Ser	Gly	Glu	Pro 10	Gln	Asn	Lys	Arg	Thr 15	Pro
3	Asp	Leu	Pro	Glu 20	Glu	Glu	Tyr	Val	Lys 25	Glu	Glu	Ile	Gln	Glu 30	Asņ	Glu
10	Glu	Ala	Val 35	Lys	Lys	Met	Leu	Val 40	Glu	Ala	Thr	Arg	Glu 45	Phe	Glu	Glu
	Val	Val 50	Val	Asp	Glu	Ser										
15																
	(2)	INF	ORMA!	NOIT	FOR	SEQ	ID 1	NO: 2	240:							
20				C	A) L B) T D) T	ENGT: YPE : OPOL	H: 6 ami OGY:	3 am no a lin	ino cid ear	acid		: 24	0 :	*		
25	Gln 1	Lys	Leu	Lys	Arg 5	Lys	Ala	Glu	Glu	Asp 10	Pro	Glu	Ala	Ala	Asp 15	Ser
30	Gly	Glu	Pro	Gln 20	Asn	Lys	Arg	Thr	Pro 25	Asp	Leu	Pro	Glu	Glu 30	Glu	Tyr
30	Val	Lys	Glu 35	Glu	Ile	Gln	Glu	Asn 40	Glu	Glu	Ala	Val	Lys 45	Lys	Met	Leu
35	Val	Glu 50		Thr	Arg	Glu	Phe 55	Glu	Glu	Val	Val	Val 60	Asp	Glu	Ser	
40	(2)	INF		(ENCE	CHA ENGT YPE :	RACT H: 1	ERIS .13 a	TICS minc		ds					
45			(xi)	SEQ						EQ I	D NO	: 24	1:			
	Lys 1		Met	Glu	Lys 5	Ser	Ser	Leu	Thr	Gln 10	His	Ser	Trp	Gln	Ser 15	Leu
50	Lys	Asp	Arg	Tyr 20	Leu	Lys	His	Leu	Arg 25		Gln	Glu	His	Lys 30	Tyr	Leu
55	Leu	Gly	Asp 35	Ala	Pro	Val	Ser	Pro 40		Ser	Gln	Lys	Leu 45		Arg	Lys
	Ala	Glu 50		Asp	Pro	Glu	Ala 55		. Asp	Ser	Gly	Glu 60		Gln	Asn	Lys
60 .	Arg 65		Pro	Asp	Leu	Pro 70		Glu	Glu	Tyr	Val		Glu	Glu	Ile	Gln 80

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	Glu	Asn	Glu	Glu	Ala 85	Val	Lys	Lys	Met	Leu 90	Val	Glu	Ala	Thr	Arg 95	Glu
5	Phe	Glu	Glu	Val 100	Val	Val	Asp	Glu	Ser 105	Pro	Pro	Asp	Phe	Glu 110	Ile	His
10	Ile															
	40.				500	CEO.	TD .	10 - 1	140							
15	(2)	INF	(i)	(ENCE A) L B) T D) T	CHAI ENGT YPE: OPOL	RACTI H: 1 ami OGY:	ERIS' 48 a no a lin	FICS: mino cid ear	aci		: 24:	2:			
20	Leu 1	Pro	Ser	Tyr	Asp 5	Glu	Ala	Glů	Arg	Thr	Lys	Ala	Glu	Ala	Thr 15	Ile
25	Pro	Leu	Val	Pro 20	Gly	Arg	Asp	Glu	Asp 25	Phe	Val	Gly	Arg	Asp 30	Asp	Phe
	Asp	Asp	Ala 35	Asp	Gln	Leu	Arg	Ile 40	Gly	Asn	qzA	Gly	Ile 45	Phe	Met	Leu
30	Thr	Phe 50		Met	Ala	Phe	Leu 55	Phe	Asn	Trp	Ile	Gly 60		Phe	Leu	Ser
35	Phe 65	Cys	Leu	Thr	Thr	Ser 70	Ala	Ala	Gly	Arg	Tyr 75	Gly	Ala	Ile	Ser	Gly 80
	Phe	Gly	Leu	ı Ser	Leu 85		Lys	Trp	Ile	Leu 90		Val	Arg	Phe	Ser 95	
40	Tyr	Ph∈	e Pro	Gly 100	_	Phe	Asp	Gly	Gln 105		Trp	Leu	Trp	Trp 110	Val	Phe
	Leu	Val	Let 115	ı Gly	Phe	Leu	Leu	Phe 120		Arg	Gly	Phe	125		Туг	Ala
45	Lys	Val		g Lys	: Met	Pro	Glu 135		Phe	Ser	Asn	Leu 140		Arg	The	Arg
50	Val 145		ı Phe	e Ile	2											
	(2)	IM	FORM	OITA	1 FOF	R SEC	D ID	NO:	243:							
55			(i)	SEQ	(A) (B)	LENG TYPE	TH: : am	24 a ino	mino acid	aci	ds					
60			(xi) SE					near ON: :	SEQ	ID N	0: 2	43:			

	Ala 1	Gly	Arg	Tyr	Gly 5	Ala	Ile	Ser	Gly	Phe 10	Gly	Leu	Ser	Leu	Ile 15	Lys
5	Trp	Ile	`Leu	Ile 20	Val	Arg	Phe	Ser								
1.0	(2)	INFO	ORMAS	rion	FOR	SEQ	ID 1	NO: 2	244:						•	
10			(i)	(A) L	ENGT	н: 5	1 am	ino		s					
15			(xi)	(D) T	OPOL	OGY:		ear	EQ I	D NO	: 24	4 :			
	Met 1	Lys	His	Leu	Ser 5	Ala	Trp	Asn	Phe	Thr 10	Lys	Leu	Thr	Phe	Leu 15	Gln
20	Leu	Trp	Glu	Ile 20	Phe	Glu	Gly	Ser	Val 25	Glu	Asn	Cys	Gln	Thr 30	Leu	Thr
25	Ser	Tyr	Ser 35	Lys	Leu	Gln	Ile	Lys 40	Tyr	Thr	Phe	Ser	Arg 45	Gly	Ser	Thr
	Phe	Туr 50	Ile				4									
30	(2)	INFO	ORMA'	rion	FOR	SEQ	ID I	VO: 2	245 :							
			(i) :		ENCE A) L			_			ds					
35			(xi)	(D) T	OPOL	OGY:		ear	EQ I	D NO	: 24!	5 :			
40	Phe 1	Ser	Ser	Asp	Phe 5	Arg	Thr	Ser	Pro	Trp	Glu	Ser	Arg	Arg	Val 15	Glu
	Ser	Lys	Ala	Thr 20	Ser	Ala	Arg	Cys	Gly 25	Leu	Trp	Gly	Ser	Gly 30	Pro	Arg
45	Arg	Arg	Pro 35	Ala	Ser	Gly	Met	Phe 40	Arg	Gly	Leu	Ser	Ser 45	Trp	Leu	Gly
50	Leu	Gln 50	Gln	Pro	Val	Ala	Gly 55	Gly	Gly	Gln	Pro	Asn 60	Gly	Asp	Ala	Pro
	Pro 65	Glu	Gln	Pro	Ser	Glu 70	Thr	Val	Ala	Glu	Ser 75	Ala	Glu	Glu	Glu	Leu 80
55	Gln	Gln	Ala	Gly	Asp 85	Gln	Glu	Leu	Leu	His 90	Gln	Ala	Lys	Asp	Phe 95	Gly
	Asn	Tyr	Leu	Phe 100	Asn	Phe	Ala	Ser	Ala 105	Ala	Thr	Lys	Lys	Ile 110	Thr	Glu
60	Ser	Val	Ala	Glu	Thr	Ala	Gln	Thr	Ile	Lys	Lys	Ser	Val	Glu	Glu	Gly

			115					120					125				
5	Lys	Ile 130	Asp	Gly	Ile	Ile	Asp 135	Lys	Thr	Ile	Ile	Gly 140	Asp	Phe	Gln	Lys	
J	Glu 145	Gln	Lys	Lys	Phe	Val 150	Glu	Glu	Gln	His	Thr 155	Lys	Lys	Ser	Gļu	Ala 160	
10	Ala	Val	Pro	Pro	Trp 165	Val	Asp	Thr	Asn	Asp 170	Glu	Glu	Thr	Ile	Gln 175	Gln	
	Gln	Ile	Leu	Ala 180	Leu	Ser	Ala	Asp	Lys 185	Arg	Asn	Phe	Leu	Arg 190	Asp	Pṛo	
15	Pro	Ala	Gly 195	Val	Gln	Phe	Asn	Phe 200	Asp	Phe	Asp	Gln	Met 205	Туг	Pro	Val	
20	Ala	Leu 210	Val	Met	Leu												
	(2)	INF	ORMA!	rion	FOR	SEQ	ID I	NO: 2	246:								
25				(A) L B) T D) T	ENGT YPE : OPOL	H: 4 ami OGY:	9 am no a lin	ino cid ear	acid							
30				SEQ													
	Met 1	Arg	Phe	Ala	Leu 5	Val	Pro	Lys	Leu	Val 10	Lys	Glu	Glu	Val	Phe 15	Trp	
35	Arg	Asn	Tyr	Phe 20	Tyr	Arg	Val	Ser	Leu 25	Ile	Lys	Gln	Ser	Ala 30	Gln	Leu	
	Thr	Ala	Leu 35	Ala	Ala	Gln	Gln	Gln 40	Ala	Ala	Gly	Lys	Gly 45	Gly	Glu	Glu	
40	Gln																
45	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	247:								
50			(i)	(A) I	ENGT	H: 7	ERIS 6 am ino a : lir	nino ncid		ls						
			(xi)	SEQ						EQ I	D NO): 24	17:				
55	Ser 1		Ser	Pro	Gly 5		Ser	Glu	Phe	Val 10		Asp	Ala	. Phe	Asp 15	Ala	
	Cys	Asn	Leu	Asn 20		Glu	Asp	Leu	Arg 25		Glu	Met	Glu	Gln 30		Val	
60	Leu	Asp	Lys 35		Gln	Glu	Glu	Thr		Val	. Leu	Glu	Glu 45		Ser	Ala	

	Asp	Trp 50	Glu	Lys	Glu	Leu	Gln 55	Gln	Glu	Leu	Gln	Glu 60	Tyr	Glu	Val	Val
5	Thr 65	Glu	Ser	Glu	Lys	Arg 70	Asp	Glu	Asn	Trp	Asp 75	Lys				
10	(2)	INF	ORMA!	rion	FOR	SEQ	ID I	VO: 2	248:							
			(i)	(A) L	ENGT	н: 6	ERIS' 2 am no a	ino		s					
15			(xi)					lin PTIO		EQ I	D NO	: 24	B:			
20	Ser 1	Pro	Trp	Glu	Ser 5	Arg	Arg	Val	Glu	Ser 10	Lys	Ala	Thr	Ser	Ala 15	Arg
	Cys	Gly	Leu	Trp 20	Gly	Ser	Gly	Pro	Arg 25	Arg	Arg	Pro	Ala	Ser 30	Gly	Met
25	Phe	Arg	Gly 35	Leu	Ser	Ser	Trp	Leu 40	Gly	Leu	Gln	Gln	Pro 45	Val	Ala	Gly
	Gly	Gly 50	Gln	Pro	Asn	Gly	Asp 55	Ala	Pro	Pro	Glu	Gln 60	Pro	Ser		
30																
	(2)	INF	ORMA'					NO:								
35				(A) I B) I D) I	ENGT YPE : OPOL	H: 6 ami OGY:	55 am no a lin	uino cid ear	acid		. 24	a .			
40	Pro	Val												Pro	Glu	Gln
	1				5					10					15	
45	Pro	Ser	· Glu	Thr 20	Val	Ala	Glu	Ser	Ala 25	Glu	Glu	Glu	Leu	Gln 30	Gln	Ala
	Gly	Asp	Gln 35		Leu	Leu	His	Gln 40		. Lys	qaA	Phe	Gly 45		Tyr	Leu
50	Phe	Asn 50		Ala	Ser	Ala	Ala 55		Lys	Lys	Ile	Thr 60		Ser	Val	Ala
	Glu 65															
55																
	(2)	INF	ORMA	MOIT	FOR	SEQ) ID	NO:	250:							
60			(i)					reris 72 ar			ds					

(A) LENGTH: 72 amino acids

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								no a lin								
			(xi)	SEQ						EQ I	D NO	: 25	0:			
5	Phe 1	Gln	Lys	Glu	Gln 5	Lys	Lys	Phe	Val	Glu 10	Glu	Gln	His	Thr	Lys 15	Lys
10	Ser	Glu	Ala	Ala 20	Val	Pro	Pro	Trp	Val 25	Asp	Thr	Asn	Asp	Glu 30	Glu	Thr
10	Ile	Gln	Gln 35	Gln	Ile	Leu	Ala	Leu 40	Ser	Ala	Asp	Lys	Arg 45	Asn	Phe	Leu
15	Arg	Asp 50	Pro	Pro	Ala	Gly	Val 55	Gln	Phe	Asn	Phe	Asp 60	Phe	Asp	Gln	Met
	Тут 65	Pro	Val	Ala	Leu	Val 70	Met	Leu								
20																
	(2)	INF	ORMA'	TION	FOR	SEQ	ID	NO:	251:							
25			7.	(A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami OGY:	8 am no a lin	ino cid ear	acid		o: 25	1:			
30	Pro 1	Phe		Cys		Ala					Ser			Phe	Ser 15	Ser
35	Pro	Ile	Leu	Ala 20	Arg	Lys	Leu	Cys	Glu 25		Ala	Ala				
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	252 :							
40			(i)		(A) I (B) T	ENGT	H: I	33 ar ino a	nino acid		ds					
45			(xi)	SEÇ				: lir [PTIC		SEQ I	D NO): 25	52:			
43	Lys 1		Asp	Pro	Ala 5		Thr	Val	Туг	Ser 10		Val	. Glu	ılle	Pro	
50	Lys	Met	: Glu	Asn 20		His	s Ser	Leu	Let 25		Met	Pro	Asp	Thr 30		Arg
	Leu	L													•	
55		•														
	(2)	INE	ORMA	4OITA	FOF	R SEQ) ID	NO:	253	:						
60			(i)	SEQ				TERI: 227			ids					

					B) T											
			(xi)		D) TO JENCI					EQ II	ON C	: 25	3:			
5																
5	Ala 1	Ser	Ala	Val	Leu 5	Leu	Asp	Leu	Pro	Asn 10	Ser	Gly	Gly	Glu	Ala 15	Gln
10	Ala	Lys	Lys	Leu 20	Gly	Asn	Asn	Cys	Val 25	Phe	Ala	Pro	Ala	Asp 30	Val	Thr
10	Ser	Glu	Lys 35	Asp	Val	Gln	Thr	Ala 40	Leu	Ala	Leu	Ala	Lys 45	Gly	Lys	Phe
15	Gly	Arg 50	Val	Asp	Val	Ala	Val 55	Asn	Cys	Ala	Gly	Ile 60	Ala	Val	Ala	Ser
	Lys 65	Thr	Тут	Asn	Leu	Lys 70	Lys	Gly	Gln	Thr	His 75	Thr	Leu	Glu	Asp	Phe .80
20	Gln	Arg	Val	Leu	Asp 85	Val	Asn	Leu	Met	Gly 90	Thr	Phe	Asn	Val	Ile 95	Arg
25	Leu	Val	Ala	Gly 100	Glu	Met	Gly	Gln	Asn 105	Glu	Pro	Asp	Gln	Gly 110	Gly	Gln
	Arg	Gly	Val 115	Ile	Ile	Asn	Thr	Ala 120	Ser	Val	Ala	Ala	Phe 125	Glu	Gly	Gln
30	Val	Gly 130	Gln	Ala	Ala	Tyr	Ser 135	Ala	Ser	Lys	Gly	Gly 140	Ile	Val	Gly	Met
	Thr 145	Leu	Pro	Ile	Ala	Arg 150	Asp	Leu	Ala	Pro	Ile 155	Gly	Ile	Arg	Val	Met 160
35	Thr	Ile	Ala	Pro	Gly 165	Leu	Phe	Gly	Thr	Pro 170	Leu	Leu	Thr	Ser	Leu 175	Pro
40	Glu	Lys	Val	Cys 180	Asn	Phe	Leu	Ala	Ser 185	Gln	Val	Pro	Phe	Pro 190	Ser	Arg
	Leu	Gly	Asp 195	Pro	Ala	Glu	Tyr	Ala 200	His	Leu	Val	Gln	Ala 205	Ile	Ile	Glu
45	Asn	Pro 210	Phe	Leu	Asn	Gly	Glu 215	Val	Ile	Arg	Leu	Asp 220	Gly	Ala	Ile	Arg
	Met 225	Gln	Pro													
50																
	(2)	INF	ORMA	rion	FOR	SEQ	ID 1	NO: 3	254 :							
55			(i)	(ENCE A) L B) T D) T	ENGT YPE:	H: 2 ami	9 am no a	ino cid		ls					
			(xi)		UENC					EQ I	D NO	: 25	4:			
60	Ser	Val	Ala	Ala	Phe	Glu	Gly	Gln	Val	Gly	Gln	Ala	Ala	Tyr	Ser	Ala

	1				5					10					15	
5	Ser	Lys	Gly	Gly 20	Ile	Val	Gly	Met	Thr 25	Leu	Pro	Ile	Ala			
10	(2)	INF		SEQU!	FOR ENCE A) L	CHA ENGT	RACT H: 6	ERIS 1 am	TICS ino		s				•	
15	Ala	Arg		(SEQ	B) T D) T UENC Gly	OPOL E DE	OGY: SCRI	lin PTIO	ear N: S					Cys	Arg	Trp
20	1 Ala	Gln	Lys	His 20	5 Lys	Asn	Trp	Arg	Phe 25	10 Gln	Lys	Thr	Arg	Gln 30	15 Thr	Trp
	Leu	Leu	Leu 35	His	Met	Tyr	Asp	Ser 40	Asp	Lys	, Val	Pro	Asp 45		His	Phe
25	Ser	Thr 50	Leu	Leu	Ala	Tyr	Leu 55	Glu	Gly	Leu	Gln	Gly 60	Arg			
30	(2)			SEQUI	FOR ENCE A) L	CHAI	RACT	ERIS	rics		c	·				
35			(xi)	(B) T D) T UENCI	YPE: OPOL	ami OGY:	no a lin	cid ear			: 25	б:			
40	1				Trp 5		Ile	Asn	Ala	Ala 10	Thr	Leu	Ser	Gln	Phe 15	Туr
,,	116	ASII	Буѕ	20	Cys	Pne										
45	(2)				FOR ENCE											
50				(A) L B) T D) T UENCI	ENGT: YPE: OPOL	H: 2 ami OGY:	2 am no a lin	ino cid ear	acid	-	: 25	7:			
55	1				Tyr 5		Leu	Thr	Leu	Met 10	Gln	Asn	Ala	Gln	Leu 15	Ser
			•	20	-	-4										

	(2) INFORMATION FOR SEQ ID NO: 258:
5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 258:
10	Lys Val Ser Tyr Leu Arg Pro Leu Asp Phe Glu Glu Ala Arg Glu Leu 1 5 10 15
	Phe Leu Cly Gln His Tyr Val Phe 20 25
15	
	(2) INFORMATION FOR SEQ ID NO: 259:
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 259:
25	Met Glu Arg Arg Cys Lys Met His Lys Arg Xaa Ile Ala Met Leu Glu 1 5 10 15
30	Pro Leu Thr Val Asp Leu Asn Pro Gln 20 25
	(2) INFORMATION FOR SEQ ID NO: 260:
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid
40	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 260:
.0	Ser His Ile Val Lys Lys-Ile Asn Asn Leu Asn Lys Ser Ala Leu Lys 1 10 15
45	Tyr Tyr Gln Leu Phe Leu Asp 20
50	(2) INFORMATION FOR SEQ ID NO: 261: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 64 amino acids
	(B) TYPE: amino acid
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 261:
	Phe Thr His Leu Ser Thr Cys Leu Leu Ser Leu Leu Leu Val Arg Met 1 5 10 15
60	Ser Gly Phe Leu Leu Ala Arg Ala Ser Pro Ser Ile Cys Ala Leu

			20					25					30		
5	Asp Se	er Ser 35	Cys	Phe	Val	Gln	Glu 40	Tyr	Cys	Ser	Ser	Туг 45	Ser	Ser	Ser
J	Cys Ph	ne Leu 50	His	Gln	His	Phe 55	Pro	Ser	Leu	Leu	Asp 60	His	Leu	Cys •	Gln
10															
15	(2) IN	VFORMA:	SEQUI	ENCE	CHA	RACT	ERI <i>S'</i>								
20		(xi)	(B) T D) T	YPE: OPOL	ami OGY:	no a lin	cid ear			: 26	2 :			
	Phe Le	eu Leu	Leu	Ala 5	Arg	Ala	Ser	Pro	Ser 10	Ile	Cys	Ala	Leu	Asp 15	Ser
25	Ser Cy	ys Phe	Val 20	Gln	Glu	Tyr									
30	(2) II	nforma'	TION	FOR	SEQ	ID	NO: 3	263:							
35			(A) L B) T D) T	ENGT YPE : OPOL	H: 5 ami OGY:	3 am no a lin	ino cid ear	acid		: 26	3 :			
40	Pro A	sp Gly	Arg	Val 5	Thr	Asn	Ile	Pro	Gln 10	Gly	Met	Val	Thr	Asp 15	Gln
	Phe G	ly Met	Ile 20	Gly	Leu	Leu	Thr	Phe 25	Ile	Arg	Ala	Ala	Glu 30	Thr	Asp
45	Pro G	ly Met 35		His	Leu	Ala	Leu 40		Ser	Asp	Leu	Thr 45		Leu	Gly
		sn Leu 50	Asn	Ser											
50														•	
	(2) I	NFORMA	MOIT	FOR	SEÇ	ID	NO:	264:							
55				(A) I (B) 1 (D) 1	LENG TYPE TOPO	TH: : am LOGY	41 ar ino a : lir	mino acid near	acio		D: 2	64:			
60	C1 *	to		n	. m		. 11% ~~	· 175 ~	· Mat		. 01	- 63			• .

10 15 Gln Leu Leu Ala Ala Val Glu Leu Phe Asn Arg Asp Trp Arg Tyr His 30 5 Lys Glu Glu Arg Val Trp Ile Thr Arg 10 (2) INFORMATION FOR SEQ ID NO: 265: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 amino acids 15 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 265: Val His Leu Ala Leu Gly Ser Asp Leu Thr Thr Leu Gly Leu Asn Leu 20 5 Asn Ser Pro Glu Asn Leu Tyr Pro 20 25 (2) INFORMATION FOR SEQ ID NO: 266: (i) SEQUENCE CHARACTERISTICS: 30 (A) LENGTH: 41 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 266: 35 His Asn Glu Asp Phe Pro Ala Leu Pro Gly Ser 1 5 40 (2) INFORMATION FOR SEQ ID NO: 267: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 75 amino acids (B) TYPE: amino acid 45 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 267: Gly Arg Ile Ile Asp Thr Ser Leu Thr Arg Asp Pro Leu Val Ile Glu 10 50 Leu Gly Gln Lys Gln Val Ile Pro Gly Leu Glu Gln Ser Leu Leu Asp Met Cys Val Gly Glu Lys Arg Arg Ala Ile Ile Pro Ser His Leu Ala 55 Tyr Gly Lys Arg Gly Phe Pro Pro Ser Val Pro Ala Asp Ala Val Val 60 Cln Tyr Asp Val Glu Leu Ile Ala Leu Ile Arg

60

349

65 70 75 5 (2) INFORMATION FOR SEQ ID NO: 268: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 268: Ile His Tyr Thr Gly Ser Leu Val Asp Gly Arg Ile Ile Asp Thr Ser 1 5 10 15 20 (2) INFORMATION FOR SEQ ID NO: 269: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 269: Cys Glu Ser Pro Glu Ser Pro Ala Gln Pro Ser Gly Ser Ser Leu Pro 30 1 5 Ala Trp Tyr His 20 35 (2) INFORMATION FOR SEQ ID NO: 270: (i) SEQUENCE CHARACTERISTICS: 40 (A) LENGTH: 95 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 270: 45 Glu Glu Ala Gly Ala Gly Arg Cys Ser His Gly Gly Ala Arg Pro 1 5 Ala Gly Leu Gly Asn Glu Gly Leu Gly Leu Gly Gly Asp Pro Asp His 25 50 Thr Asp Thr Gly Ser Arg Ser Lys Gln Arg Ile Asn Asn Trp Lys Glu 35 40 Ser Lys His Lys Val Ile Met Ala Ser Ala Ser Ala Arg Gly Asn Gln 55 Asp Lys Asp Ala His Phe Pro Pro Pro Ser Lys Gln Ser Leu Leu Phe

70

Cys Pro Lys Ser Lys Leu His Ile His Arg Ala Glu Ile Ser Lys

350

95

90

5 (2) INFORMATION FOR SEQ ID NO: 271: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 271: Ser Lys Gln Arg Ile Asn Asn Trp Lys Glu Ser Lys His Lys Val Ile 10 15 Met Ala Ser Ala Ser Ala Arg 20 20 (2) INFORMATION FOR SEQ ID NO: 272: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 32 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 272: Leu Phe His Trp Ala Cys Leu Asn Glu Arg Ala Ala Gln Leu Pro Arg 30 Asn Thr Ala Xaa Ala Gly Tyr Gln Cys Pro Ser Cys Asn Gly Pro Ser 25 35 40 (2) INFORMATION FOR SEQ ID NO: 273: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 185 amino acids (B) TYPE: amino acid 45 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 273: Phe Tyr Ile Tyr Tyr Arg Pro Thr Asp Ser Asp Asn Asp Ser Asp Tyr 50 Lys Lys Asp Met Val Glu Gly Asp Lys Tyr Trp His Ser Ile Ser His Leu Gln Pro Glu Thr Ser Tyr Asp Ile Lys Met Gln Cys Phe Asn Glu 55 40 Gly Gly Glu Ser Glu Phe Ser Asn Val Met Ile Cys Glu Thr Lys Ala 55 s 60

Arg Lys Ser Ser Gly Gln Pro Gly Arg Leu Pro Pro Pro Thr Leu Ala

WO 98/42738 PCT/US98/05311

	65					70					75					80
5	Pro	Pro	Gln	Pro	Pro 85	Leu	Pro	Glu	Thr	Ile 90	Glu	Arg	Pro	Val	Gly 95	Thr
J	Gly	Ala	Met	Val 100	Ala	Arg	Ser	Ser	Asp 105	Leu	Pro	Tyr	Leu	Ile 110	Val	Gly
10	Val	Val	Leu 115	Gly	Ser	Ile	Val	Leu 120	Ile	Ile	Val	Thr	Phe 125	Ile	Pro	Phe
	Cys	Leu 130	Trp	Arg	Ala	Trp	Ser 135	Lys	Gln	Lys	His	Thr 140	Thr	Asp	Leu	Gļy
15	Phe 145	Pro	Arg	Ser	Ala	Leu 150	Pro	Pro	Ser	Cys	Pro 155	Tyr	Thr	Met	Val	Pro 160
20	Leu	Gly	Gly	Leu	Pro 165	Gly	His	Gln	Ala	Val 170	Asp	Ser	Pro	Thr	Ser 175	Val
	Ala	Ser	Val	Asp 180	Gly	Pro	Val	Leu	Met 185							
25	(2)	INF	ORMAC	rion	FOR	SEQ	ID I	NO: 2	274:							
			(i)	SEQU	ENCE	CHA	RACT	ERIS'	TICS	:						
30			(xi)	(B) T D) T	YPE : OPOL	ami : OGY	6 am no a lin PTIO	cid ear			: 27	4:			
35	Tyr 1	Ile	Tyr	Tyr	Arg 5	Pro	Thr	Asp	Ser	Asp	Asn	Asp	Ser	Asp	Туг 15	Lys
	Lys	Asp	Met	Val 20	Glu	Gly	Asp	Lys	Туг 25	Trp	His	Ser	Ile	Ser 30	His	Leu
40	Gìn	Pro	Glu 35	Thr	Ser	Tyr	Asp	Ile 40	Lys	Met	Gln	Cys	Phe 45	Asn	Glu	Gly
45	Gly	Glu 50		Glu	Phe	Ser	Asn 55	Val	Met	Ile	Cys	Glu 60		Lys	Ala	Arg
73	Lys 65	Ser														
50	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	275 :							
55				,	(A) I (B) 1 (D) 1	LENGT TYPE TOPOI	H: : : am: : : COGY	TERIS 30 ar ino a : lir	nino acid near	acio		D: 27	75 :			
60	Asn 1		Arg	Ala	Leu 5		His	: Arg	Met	Pro		ı Pro	Pro	Lys	: Ile	

	Thr	Ala	Lys	Phe 20	Asn	Asn	Asn	Lys	Arg 25	Lys	Asn	Leu	Ser	Leu 30		
5																
	(2)	INF	ORMAT	NOI	FOR	SEQ	ID N	10: 2	276 :						•	
10			(i) :	(A) L B) T D) T	ENGT YPE : OPOL	H: 1 ami OGY:	85 a no a lin	mino cid ear	aci		: 27	6 :			a.
15	Asn 1	Thr	Asn	Gln	Arg 5	Glu	Ala	Leu	Gln	Tyr 10	Ala	Lys	Asn	Phe	Gln 15	Pro
20	Phe	Ala	Leu	Asn 20	His	Gln	Lys	Asp	Ile 25	Gln	Val	Leu	Met	Gly 30	Ser	Leu
20	Val	Tyr	Leu 35	Arg	Gln	Gly	Ile	Glu 40	Asn	Ser	Pro	Tyr	Val 45	His	Leu	Leu
25	Asp	Ala 50	Asn	Gln	Trp	Ala	Asp 55	Ile	Cys	Asp	Ile	Phe 60	Thr	Arg	qaA	Ala
	Суs 65	Ala	Leu	Leu	Gly	Leu 70	Ser	Val	Glu	Ser	Pro 75	Leu	Ser	Val	Ser	Phe 80
30	Ser	Ala	Gly	Cys	Val 85	Ala	Leu	Pro	Ala	Leu 90	Ile	Asn	Ile	Lys	Ala 95	Val
35	Ile	Glu	Gln	Arg 100	Gln	Cys	Thr	Gly	Val 105	Trp	Asn	Gln	Lys	Asp 110	Glu	Leu
	Pro	Ile	Glu 115	Val	Asp	Leu	Gly	Lys 120	Lys	Cys	Trp	Tyr	His 125	Ser	Ile	Phe
40	Ala	Cys 130	Pro	Ile	Leu	Arg	Gln 135	Gln	Thr	Thr	Asp	Asn 140	Asn	Pro	Pro	Met
	Lys 145		Val	Cys	Gly	His 150		Ile	Ser	Arg	Asp 155	Ala	Leu	Asn	Lys	Met 160
45	Phe	Asn	Gly	Ser	Lys 165		Lys	Cys	Pro	Tyr 170	Cys	Pro	Met	Glu	Gln 175	Ser
50	Pro	Gly	Asp	Ala 180	-	Gln	Ile	Phe	Phe 185							
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	277 :							
55			(i)		ENCE (A) I (B) 5	LENGT TYPE :	TH: 6	55 an ino a	nino acid		ls					
			(xi)		UENC					SEQ I	D NO): 27	77:			

	Ser 1	Tyr	Leu	Ser	Ala 5	Cys	Phe	Ala	Gly	Cys 10	Asn	Ser	Thr	Asn	Leu 15	Thr
5	Gly	Cys	Ala	Cys 20	Leu	Thr	Thr	Val	Pro 25	Ala	Glu	Asn	Ala	Thr 30	Val	Val
	Pro	Gly	Lys 35	Суѕ	Pro	Ser	Pro	Gly 40	Cys	Gln	Glu	Ala	Phe 45	Leu	Thr	Phe
10	Leu	Cys 50	Val	Met	Cys	Ile	Cys 55	Ser	Leu	Ile	Gly	Ala 60	Met	Ala	Arg	His
15	Pro 65															
20	(2)	INF	ORMA	SEQU))	ENCE A) L B) T	CHA ENGT YPE:	RACT H: 8 ami		TICS ino cid		ls					
25	Pro		(xi) Val							_	Ser	: 27		Leu	Lys 15	Ser
30			ı Leu	Gly 20	Val	Leu	Phe	Leu	Leu 25		. Arg	Leu	Leu	Gly 30	Phe	Ile
	Pro	Pro	Pro 35		Ile	Phe	Gly	Ala 40	Gly	Ile	Asp	Ser	Thr 45	-	Leu	Phe
35	Тг	Ser 50		Phe	Cys	Gly	Glu 55		Gly	Ala	. Cys	Val 60	Leu	Tyr	Asp	Asn
40	69	5	_			Leu 70		Val	Ser	Ile	2 Ala 75		Ala	Leu	Lys	Ser 80
	Phe	≥ Ala	a Phe	e Ile	:											
45	(2) IN	FORM	4OITA	1 FOF	SEQ) ID	NO:	279 :							
50				SEQU	(A) (B) ((D)	LENG' TYPE TOPO!	TH: : am LOGY	182 ino : li	amin acid near	o ac		O: 27	79:			
55		n Se 1	r Le	u Phe		c Arg	g Ph	e Vai	l Arg	y Va 1		y Val	Pro	o Thi	c Va 1	l Asp 5
	Le	u As	p Al	a Gli 20		/ Arg	g Al	a Ar	g Ala 2		r Le	u Cys	s Xa	a Xaa	_	r Asn
60	Tr	p Ar	g Ty	r Ly:	s Ası	n Lev	G1 د	y Ası	n Lei	u Pr	o Hi	s Vai	1 G1	n Le	u Le	u Pro

			2.5					40								
			35					40					45			
5	Glu	Phe 50	Ser	Thr	Ala	Asn	Ala 55	Gly	Leu	Leu	Tyr	Asp 00	Phe	Gln	Leu	Ile
•	Asn 65	Val	Glu	Asp	Phe	Gln 70	Gly	Val	Gly	Glu	Ser 75	Glu	Pro	Asn	Pro	Туг 80
10	Phe	Tyr	Gln	Asn	Leu 85	Gly	Glu	Ala	Glu	Tyr 90	Val	Val	Ala	Leu	Phe 95	Met
	Tyr	Met	Cys	Leu 100	Leu	Gly	Tyr	Pro	Ala 105	Asp	Lys	Ile	Ser	Ile 110	Leu	Thr
15	Thr	Tyr	Asn 115	Gly	Gln	Lys	His	Leu 120	Ile	Arg	Asp	Ile	Ile 125	Asn	Arģ	Arg
20	Cys	Gly 130	Asn	Asn	Pro	Leu	Ile 135	Gly	Arg	Pro	Asn	Lys 140	Val	Thr	Thr	Val
20	Asp 145	Arg	Phe	Gln	Gly	Gln 150	Gln	Asn	Asp	Туг	Ile 155	Leu	'Leu	Ser	Leu	Val 160
25	Arg	Thr	Arg	Ala	Val 165	Gly	His	Leu	Arg	Asp 170	Val	Arg	Arg	Leu	Val 175	Val
	Ala	Met	Ser	Arg 180	Ala	Arg										
30																
	(2)	INF	ORMA!	NOI	FOR	SEQ	ID I	O: 1	280:							
35				(A) L B) T D) T	ENGT YPE : OPOL	H: 7 ami OGY:	ERIS' 7 am no a lin PTIO	ino cid ear	acid		: 28	0 :			
40	Leu 1	Val	Lys	Glu	Ala 5	Lys	Ile	Ile	Ala	Met	Thr	Cys	Thr	His	Ala 15	Ala
		Lys	Arg	His		Leu	Val	Lys	Leu 25		Phe	Lys	Tyr	Asp 30		Ile
45	Leu	Met	Glu 35		Ala	Ala	Gln	Ile 40		Glu	Ile	Glu	Thr 45		Ile	Pro
50	Leu	Leu 50		Gln	Asn	Pro	Gln 55		Gly	Phe	Ser	Arg 60		Lys	Arg	Trp
	Ile 65	Met	Ile	Gly	Asp	His 70	His	Gln	Leu	Pro	Pro 75	Val	Ile			
55																
	(2)	INF	ORMA'	NOIT	FOR	SEQ	ID	NO:	281:							
60			(i)					ERIS			ds					

			.	(D) T	OPOL	OGY:	no a lin	ear		_					
5	_			SEQ												
5	Asp 1	Thr	Tyr	Pro	Asn 5	Glu	Glu	Lys	Gln	Gln 10	Glu	Arg	Val	Phe	Pro 15	Xaa
10	Xaa	Ser	Ala	Met 20	Val	Asn	Asn	Gly	Ser 25	Leu	Ser	Tyr	Asp	His 30	Glu	Arg
	Asp	Gly	Arg 35	Pro	Thr	Glu	Leu	Gly 40	Gly	Cys	Xaa	Ala	Ile 45	Val	Arg	Asr
15	Leu	His 50	Tyr	Asp	Thr	Phe	Leu 55	Val	Ile	Arg	Tyr	Val 60	Lys	Arg	His	Leu
	Thr 65	Ile	Met	Met	Asp	Ile 70	Asp	Gly	Lys	His	Glu 75	Trp	Arg	Asp	Cys	11e
20	Glu	Val	Pro	Gly	Val 85	Arg	Leu	Pro	Arg	Gly 90	Tyr	Tyr	Phe	Gly	Thr 95	Ser
25	Ser	Ile	Thr	Gly 100	Asp	Leu	Ser	Asp	Asn 105	His	Asp	Val	Ile	Ser 110	Leu	Lys
	Leu	Phe	Glu 115	Leu	Thr	Val	Glu	Arg 120	Thr	Pro	Glu	Glu	Glu 125			
30	(2)	TMEC	ימשכר	noin	FOR	ero.	TD 1	io. i	ogo.							
	(2)			SEQUI	ENCE	CHAI	RACT	ERIS	rics							
35				(B) T D) T	YPE: OPOL	ami OGY:	no a	ear							
				SEQ												
40	Leu 1	Lys	Arg	Glu	His 5	Ser	Leu	Ser	Lys	Pro 10	Tyr	Gln	Gly	Val	Gly 15	Thr
	Gly	Ser	Ser	Ser 20	Leu	Trp	Asn	Leu	Met 25	Gly	Asn	Ala	Met	Val 30	Met	Thr
45	Gln	Tyr	Ile 35	Arg	Leu	Thr	Pro	Asp 40	Met	Gln	Ser	Lys	Gln 45	Gly	Ala	Let
50	Trp	Asn 50	Arg	Val	Pro	Cys	Phe 55	Leu	Arg	Ąsp	Trp	Glu 60	Leu	Gln	Val	His
	Phe 65	Lys	Ile	His	Gly	Gln 70	Gly	Lys	Lys	Asn	Leu 75	His	Gly	Asp	Gly	Let 80
55	Ala	Ile	Trp	Tyr	Thr 85			•								

(2) INFORMATION FOR SEQ ID NO: 283:

	(1) SEQUENCE CHARACTERISTICS: (A) LENGTH: 32 amino acids
	(B) TYPE: amino acid
5	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 283:
•	Pro Gly Thr Leu Gln Cys Ser Ala Leu His His Asp Pro Gly Cys Ala 1 5 10 15
10	Asn Cys Ser Arg Phe Cys Arg Asp Cys Ser Pro Pro Ala Cys Gln Cys 20 25 30
15	
	(2) INFORMATION FOR SEQ ID NO: 284:
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 284: Phe Leu Tyr Asp Val Leu Met Xaa His Glu Ala Val Met Arg Thr His
	1 5 10 15
30	Gln Ile Gln Leu Pro Asp Pro Glu Phe Pro Ser 20 25
35	(2) INFORMATION FOR SEQ ID NO: 285:
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 6 amino acids(B) TYPE: amino acid
40	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 285:
	Gly Trp Tyr Trp Cys Gly 1 5
45	
	(2) INFORMATION FOR SEQ ID NO: 286:
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 129 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 286:
55	Met Lys Val Gly Ala Arg Ile Arg Val Lys Met Ser Val Asn Lys Ala 1 5 10 15
60	His Pro Val Val Ser Thr His Trp Arg Trp Pro Ala Glu Trp Pro Gln 20 25 30

	Met	Phe	Leu 35	His	Leu	Ala	Gln	G1u 40	Pro	Arg	Thr	Glu	Val 45	Lys	Ser	Arg
5	Pro	Leu 50	Gly	Leu	Ala	Gly	Phe 55	Ile	Arg	Gln	Asp	Ser 60	Lys	Thr	Arg	Lys
	Pro 65	Leu	Glu	Gln	Glu	Thr 70	Ile	Met	Ser	Ala	Ala 75	Asp	Thr	Ala	Leu	Trp 80
10	Pro	Tyr	Gly	His	Gly 85	Asn	Arg	Glu	His	Glņ 90	Glu	Asn	Glu	Leu	Gln 95	Lys
15	Tyr	Leu	Gln	Tyr 100	Lys	Asp	Met	His	Leu 105	Leu	Asp	Ser	Gly	Gln 110	Ser	Leu
	Gly	His	Thr 115	His	Thr	Leu	Gln	Gly 120	Ser	His	Asn	Leu	Thr 125	Ala	Leu	Asn
20	Ile															
	(2)	INF	ORMA'	TION	FOR	SEO	ID	NO:	287:							
25	,			SEQU	ENCE	СНА	RACT		TICS		ls					
30			(xi)	(B) 1	YPE :	ami : OGY.	no a	cid ear			: 28	7:			
	Ser 1	Leu	His	Lys	Asn 5	Ser	Val	Ser	Gln	Ile 10		Val	Leu	Ser	Gly 15	Gly
35	Lys	Ala	Lys	Cys 20		Gln	Phe	Cys	Thr 25		Gly	Met	Asp	Gly 30	Gly	Met
40	Ser	Ile	Trp 35		Val	Lys	Ser	Leu 40		Ser	Ala	. Leu	Lys 45		Leu	Lys
	Ile															
45	(2)	INF	ORMA	NOITA	I FOR	SEQ) ID	NO:	288 :							
50					(A) 1 (B) '	LENG TYPE TOPO!	TH: : am LOGY	21 ar ino a : lir	mino acid near	aci		D: 21	38 :			
55	Glu 1		. Ser	Lys	S Ser		His	s Alá	a Gly	/ Let) Lev	ı Phe	e Ser	Va]	l Ala

(2) INFORMATION FOR SEQ ID NO: 289:

5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 289:
10	Tyr Met Gly Lys Gly Ser Met Thr Gly Leu Ala Leu Lys His Met Phe 1 5 10 15
15	Glu Arg Ser Phe Thr 20
	(2) INFORMATION FOR SEQ ID NO: 290:
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 27 amino acids(B) TYPE: amino acid(D) TOPOLOGY: linear
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 290:
	Val Thr Gly Ile Ile Asp Ser Leu Thr Ile Ser Pro Lys Ala Ala Arg 1 5 10 15
30	Val Gly Leu Gln Tyr Ser Thr Gln Val His 20 25
35	(2) INFORMATION FOR SEQ ID NO: 291: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 amino acids (B) TYPE: amino acid
40	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 291:
	Thr Glu Phe Thr Leu Arg Asn Phe Asn Ser Ala Lys Asp Met Lys Lys 1 5 10 15
45	Ala Val Ala His Met Lys Tyr Met 20
50	(2) INFORMATION FOR SEQ ID NO: 292:
55	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 292:
60	Gly Lys Gly Ser Met Thr Gly Leu Ala Leu Lys His Met Phe Glu Arg 1 10 15

PCT/US98/05311

60

Ser Phe Thr Gln Gly Glu Gly Ala Arg Pro Phe 5 (2) INFORMATION FOR SEQ ID NO: 293: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 44 amino acids 10 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 293: Ser Thr Arg Val Pro Arg Ala Ala Ile Val Phe Thr Asp Gly Arg Ala 15 5 Gln Asp Asp Val Ser Glu Trp Ala Ser Lys Ala Lys Ala Asn Gly Ile 25 Thr Met Tyr Ala Val Gly Val Gly Lys Ala Ile Glu 20 35 25 (2) INFORMATION FOR SEQ ID NO: 294: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 42 amino acids (B) TYPE: amino acid 30 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 294: Glu Glu Leu Gln Glu Ile Ala Ser Glu Pro Thr Asn Lys His Leu Phe 1 5 35 Tyr Ala Glu Asp Phe Ser Thr Met Asp Glu Ile Ser Glu Lys Leu Lys 20 Lys Gly Ile Cys Glu Ala Leu Glu Asp Ser 40 (2) INFORMATION FOR SEQ ID NO: 295: 45 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 11 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 295: Thr Gln Arg Leu Glu Glu Met Thr Gln Arg Met 5 55 (2) INFORMATION FOR SEQ ID NO: 296: (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 10 amino acids

	(xi		PE: amin POLOGY: DESCRIP	linear	EQ ID NO:	, 296:	
5	Pro Gln Gl 1	y Cys Pro 5	Glu Gln	Pro Leu	His 10		
10	(2) INFORM	NATION FOR	SEQ ID N	0: 297:			
15		(B) T	ENGTH: 33 PE: amir PPOLOGY:	amino a no acid linear	acids	297:	
20	Arg Cys Ly l	s Lys Cys 5	Thr Glu	Gly Pro	Ile Asp 10	Leu Val	Phe Val Ile 15
20	Asp Gly Se	er Lys Ser 20	Leu Gly	Glu Glu 25	Asn Phe	Glu Val	Val Lys Gln
25	Phe						
30			CHARACTE	RISTICS			
		(B) T	/PE: amir	no acid			
35	(x)		OPOLOGY:	linear	EQ ID NO	: 298:	
35		(D) TO	OPOLOGY: E DESCRIF	linear TION: SE			Leu Arg Ala 15
35 40	Met Ala Al 1	(D) TO i) SEQUENCE .a Leu Leu 5	DPOLOGY: DESCRIF	linear TION: SE	Gly Arg	His Cys	-
	Met Ala Al 1 His Phe Se Thr Ala Ly	(D) TO i) SEQUENCE La Leu Leu 5 er Pro Gln 20 vs Glu Glu	DPOLOGY: DESCRIF Leu Arg Leu Cys Met Glu	linear PTION: SE His Val Ile Arg 25 Arg Phe 40	Gly Arg 10 Asn Ala Trp Asn	His Cys Val Pro Lys Asn 45	15 Leu Gly Thr
40	Met Ala Al 1 His Phe Se Thr Ala Ly	(D) TO i) SEQUENCE La Leu Leu 5 er Pro Gln 20 vs Glu Glu	DPOLOGY: DESCRIF Leu Arg Leu Cys Met Glu	linear PTION: SE His Val Ile Arg 25 Arg Phe 40	Gly Arg 10 Asn Ala Trp Asn	His Cys Val Pro Lys Asn 45	15 Leu Gly Thr 30
40	Met Ala Al 1 His Phe Se Thr Ala Ly 3 Asn Arg Pr	(D) TO i) SEQUENCE La Leu Leu 5 er Pro Gln 20 vs Glu Glu	DPOLOGY: DESCRIF Leu Arg Leu Cys Met Glu Pro His 55	linear PTION: SE His Val Ile Arg 25 Arg Phe 40 Ile Thr	Gly Arg 10 Asn Ala Trp Asn	His Cys Val Pro Lys Asn 45 Ser	15 Leu Gly Thr 30
40	Met Ala Al 1 His Phe Se Thr Ala Ly 3 Asn Arg Pr 50 (2) INFORM	(D) TO i) SEQUENCE La Leu Leu 5 er Pro Gln 20 vs Glu Glu 35 co Leu Ser MATION FOR) SEQUENCE (A) Li (B) T	DPOLOGY: DESCRIF DESCRIF Leu Arg Leu Cys Met Glu Pro His 55 SEQ ID N CHARACTE ENGTH: 3: YPE: amin DPOLOGY:	linear PTION: SE His Val Ile Arg 25 Arg Phe 40 Ile Thr IO: 299: ERISTICS 2 amino no acid linear	Gly Arg 10 Asn Ala Trp Asn Ile Tyr : acids	His Cys Val Pro Lys Asn 45 Ser 60	15 Leu Gly Thr 30

Trp Asp Leu Gly Lys Gly Leu Lys Ile Pro Gln Leu Tyr Gln Ser Gly 20 5 10 (2) INFORMATION FOR SEQ ID NO: 300: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 amino acids (B) TYPE: amino acid 15 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 300: Met Ala Ala Leu Leu Leu Arg His Val Gly Arg His Cys Leu Arg Ala 5 20 His 25 (2) INFORMATION FOR SEQ ID NO: 301: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 amino acids 30 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 301: Val Lys Ser Leu Cys Leu Gly Pro Ala Leu Ile His Thr Ala Lys Phe 35 5 10 Ala Leu 40 (2) INFORMATION FOR SEQ ID NO: 302: (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 23 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 302: 50 Val Phe Pro Leu Met Tyr His Thr Trp Asn Gly Ile Arg His Leu Met Trp Asp Leu Gly Lys Gly Leu 55 (2) INFORMATION FOR SEQ ID NO: 303: 60

(i) SEQUENCE CHARACTERISTICS:

```
(A) LENGTH: 22 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 303:
 5
     Arg Val Trp Asp Val Arg Pro Phe Ala Pro Lys Glu Arg Cys Val Lys
             5
      Ile Phe Gln Gly Asn Val
10
                  20
      (2) INFORMATION FOR SEQ ID NO: 304:
15
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 30 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
20
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 304:
     His Asn Phe Glu Lys Asn Leu Leu Arg Cys Ser Trp Ser Pro Asp Gly
       1
                     5
                                         10
25
      Ser Lys Ile Ala Ala Gly Ser Ala Asp Arg Phe Val Tyr Val
                                     25
                  20
30
      (2) INFORMATION FOR SEQ ID NO: 305:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 30 amino acids
                    (B) TYPE: amino acid
35
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 305:
      Trp Asp Thr Thr Ser Arg Arg Ile Leu Tyr Lys Leu Pro Gly His Ala
             5
40
      Gly Ser Ile Asn Glu Val Ala Phe His Pro Asp Glu Pro Ile
                  20
                                    25
45
      (2) INFORMATION FOR SEQ ID NO: 306:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 20 amino acids
50
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 306:
     Val Arg Gly Arg Thr Val Leu Arg Pro Gly Leu Asp Ala Glu Pro Glu
55
      1 5
                          10
      Leu Ser Pro Glu
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(2) INFORMATION FOR SEQ ID NO: 307:
             (i) SEQUENCE CHARACTERISTICS:
5
                    (A) LENGTH: 19 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 307:
10
     Glu Gln Arg Val Leu Glu Arg Lys Leu Lys Lys Glu Arg Lys Lys Glu
     Glu Arg Gln
15
      (2) INFORMATION FOR SEQ ID NO: 308:
20
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 13 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 308:
25
      Arg Leu Arg Glu Ala Gly Leu Val Ala Gln His Pro Pro
                      5
30
      (2) INFORMATION FOR SEQ ID NO: 309:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 17 amino acids
35
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 309:
      Gly Arg Ile Pro Ala Pro Ala Pro Ser Val Pro Ala Gly Pro Asp Ser
40
                                10
                        5
      Arg
45
       (2) INFORMATION FOR SEQ ID NO: 310:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 42 amino acids
 50
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 310: ..
 55
       Thr Gly Cys Val Leu Val Leu Ser Arg Asn Phe Val Gln Tyr Ala Cys
       Phe Gly Leu Phe Gly Ile Ile Ala Leu Gln Thr Ile Ala Tyr Ser Ile
                    20
 60
```

	Leu	Trp	Asp 35	Leu	Lys	Phe	Leu	Met 40	Arg	Asn						
5	(2)	INF	ORMA:	rion	FOR	SEO	I DI	NO: .	311:							
					ENCE										•	
10			(xi)	(A) L B) T D) T	ENGT YPE : OPOL	H: 5 ami OGY:	5 am no a lin	ino cid ear	acid		. 21	1 .			
	Com															
15	ser 1	Arg	Ser	GIU	G1y 5	Lys	Ser	мес	Pne	10	Gly	Val	Pro	Thr	Met 15	Arg
	Glu	Ser	Ser	Pro 20	Lys	Gln	Tyr	Met	Gln 25	Leu	Gly	Gly	Arg	Val 30	Leu	Leu
20	Val	Leu	Met 35	Phe	Met	Thr	Leu	Leu 40	His	Phe	Asp	Ala	Ser 45	Phe	Phe	Ser
25	Ile	Val 50	Gln	Asn	Ile	Val	Gly 55									
	(2)	INFO	ORMAT	rion	FOR	SEO	ID N	vo: 1	312:							
30			(i) :				•									
			(-, .	(A) L B) T D) T	ENGT YPE :	H: 6 ami	0 am no a	ino cid		s	,	•			
35			(xi)							EQ II	ON C	: 31:	2:			
55	Gly 1	Thr	Ala	Glu		Phe	Ala	qzA	Gln	Phe	Leu	Arg	Val	Thr	Lvs	
40					5					10					15	Gln
	Tyr	Leu	Pro			Ala	Arg	Leu		10		Ser	Thr	Phe 30	15	
			Pro Ile 35	His 20	Val				Cys 25	10 Leu	Ile			30	15 Leu	Glu
45	Asp	Gly	Ile	His 20 Arg	Val Met	Trp	Phe	Gln 40	Cys 25 Trp	10 Leu Ser	Ile Glu	Gln	Arg	30	15 Leu	Glu
	Asp Asp	Gly Thr 50	Ile 35 Thr	His 20 Arg Trp	Val Met Asn	Trp Cys	Phe Gly 55	Gln 40 Tyr	Cys 25 Trp Leu	10 Leu Ser	Ile Glu	Gln Ser	Arg	30	15 Leu	Glu
45 50	Asp Asp	Gly Thr 50	Ile 35	His 20 Arg Trp	Val Met Asn	Trp Cys	Phe Gly 55	Gln 40 Tyr	Cys 25 Trp Leu	10 Leu Ser	Ile Glu	Gln Ser	Arg	30	15 Leu	Glu
	Asp Asp	Gly Thr 50	Ile 35 Thr	His 20 Arg Trp Trp	Val Met Asn FOR ENCE A) L	Trp Cys SEQ CHA ENGT	Phe Gly 55 ID N RACTI H: 1	Gln 40 Tyr NO: I	Cys 25 Trp Leu 313: rics ino	Leu Ser Leu	Ile Glu Ala	Gln Ser	Arg	30	15 Leu	Glu
	Asp Asp	Gly Thr 50	Ile 35 Thr	His 20 Arg Trp Trp (((((((((((((((((((((((((((((((Val Met Asn FOR ENCE A) L B) T D) T	Trp Cys SEQ CHAL ENGT YPE: OPOL	Phe Gly 55 ID N RACTI H: 1 ami OGY:	Gln 40 Tyr NO: : ERIS' 7 am no a lin	Cys 25 Trp Leu 313: TICS ino cid ear	10 Leu Ser Leu :	Ile Glu Ala	Gln Ser 60	Arg 45	30	15 Leu	Glu
50	Asp Asp	Gly Thr 50	Ile 35 Thr ORMAT	His 20 Arg Trp Trp ((() SEQ	Val Met Asn FOR ENCE A) L B) T D) T UENCE	Trp Cys SEQ CHAL ENGT YPE: OPOL	Phe Gly 55 ID N RACTH H: 1 ami OGY: SCRI	Gln 40 Tyr NO: 1 ERIS' 7 am no a lin PTIO	Cys 25 Trp Leu 313: TICS ino cid ear N: S	10 Leu Ser Leu Leu EQ I	Ile Glu Ala S	Gln Ser 60	Arg 45	30 Asp	15 Leu Tyr	Glu

Leu

```
5
      (2) INFORMATION FOR SEQ ID NO: 314:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 8 amino acids
10
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 314:
      Leu Met Arg Asn Glu Ser Arg Ser
15
                       5
      (2) INFORMATION FOR SEQ ID NO: 315:
20
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 13 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
25
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 315:
      Ala Ser Phe Leu Leu Ser Arg Thr Ser Trp Gly Thr Ala
                        5
30
      (2) INFORMATION FOR SEQ ID NO: 316:
             (i) SEQUENCE CHARACTERISTICS:
35
                     (A) LENGTH: 20 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 316:
40
      Phe Ile Ser Phe Ala Asn Ser Arg Ser Ser Glu Asp Thr Lys Gln Met
      Met Ser Ser Phe
45
      (2) INFORMATION FOR SEO ID NO: 317:
50
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 27 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 317:
55
      Asp Pro Arg Arg Pro Asn Lys Val Leu Arg Tyr Lys Pro Pro Pro Ser
      Glu Cys Asn Pro Ala Leu Asp Asp Pro Thr Pro
60
                   20
```

5	(2) INFORMATION FOR SEQ ID NO: 318:
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 30 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 318:
	Asp Tyr Met Asn Leu Leu Gly Met Ile Phe Ser Met Cys Gly Leu Met 1 5 10 15
15	Leu Lys Leu Lys Trp Cys Ala Trp Val Ala Val Tyr Cys Ser 20 25 30
20	(2) INFORMATION FOR SEQ ID NO: 319:
25	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 319:
30	Met Leu Ser Ile Ser Ala Val Val Met Ser Tyr Leu Gln Asn Pro Glr 1 5 10 15
	Pro Met Thr Pro Pro Trp 20
35	(2) INFORMATION FOR SEQ ID NO: 320:
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 52 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 320:
45	Ala Ala Gly Asp Gly Asp Val Lys Leu Gly Thr Leu Gly Ser Gly Ser 1 5 10 15
	Glu Ser Ser Asn Asp Gly Gly Ser Glu Ser Pro Gly Asp Ala Gly Ala 20 25 30
50	Ala Ala Xaa Gly Gly Gly Trp Ala Ala Ala Ala Leu Ala Leu Leu Th 35 40 45
55	Gly Gly Glu 50
	(2) INFORMATION FOR SEQ ID NO: 321:
60	(i) SEQUENCE CHARACTERISTICS:

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				(1 (1	B) TY	PE: OPOLO	amiı XGY:	no ao line	ear							
5			(xi)	SEQ	JENCE	E DES	SCRI	PTIO	1: SE	SQ II	ОИС	: 32:	1:			
	Ala 1	Ala	Asp	Asn	Tyr 5	Gly	Ile	Pro	Arg	Ala 10	Cys	Arg	Asn	Ser	Ala 15	Arg
10	Ser	Tyr	Gly	Ala 20	Ala	Trp	Leu	Leu	Leu 25	Xaa	Pro	Ala	Gly	Ser 30	Ser	Arg
	Val	Glu	Pro 35	Thr	Gln	Asp	Ile	Ser 40	Ile	Ser	Asp	Gln	Leu 45	Gly	Gly	Gln
15	Asp	Val 50	Pro	Val	Phe	Arg	Asn 55	Leu	Ser	Leu	Leu	Val 60	Val	Gly	Val	Gly
20	Ala 65	Val	Phe	Ser	Leu	Leu 70	Phe	His	Leu	Gly	Thr 75	Arg	Glu	Arg	Arg	Arg 80
20	Pro	His	Ala	Xaa	Glu 85	Pro	Gly	Glu	His	Thr 90	Pro	Leu	Leu	Ala	Pro 95	Ala
25	Thr	Ala	Gln	Pro 100	Leu	Leu	Leu	Trp	Lys 105	His	Trp	Leu	Arg	Glu 110	Xaa	Ala
	Phe	Tyr	Gln 115	Val	Gly	Ile	Leu	Туг 120	Met	Thr	Thr	Arg	Leu 125	Ile	Val	Asn
30	Leu	Ser 130		Thr	Tyr	Met	Ala 135	Met	Tyr	Leu	Thr	Туг 140	Ser	Leu	His	Leu
35	Pro 145		Lys	Phe	Ile	Ala 150	Thr	Ile	Pro	Leu	Val 155		Tyr	Leu	Ser	Gly 160
55	Phe	Leu	Ser	Ser	Phe 165	Leu	Met	Lys	Pro	Ile 170		Lys	Cys	Ile	Gly 175	
40	Asn															
	(2)	INF	ORMA	MOIT	FOR	SEQ	ID	NO:	322:							
45			(i)	_		LENG	гн: 🗀	243 a	STICS amino acid		ids					
50			(xi)		(D) (near ON: S	SEQ :	ID N	D: 32	22:			
	Arg		e Thr	Asr	Asr S		Glu	ı Gly	/ Lys	Trp		ı Gly	y Arg	Thi	Ala 15	
55	Gl	/ Sei	г Туг	Gl ₃ 20		: Ile	e Lys	s Thi	Thr 25		a Va	l Glu	u Ile	2 Xaa		c Asp
	Sei	c Lei	ı Lys		ı Lys	s Lys	s Ası	Sei 40		r Gly	y Al	a Pro	o Se:		g Pro	o Ile
60	۰,			۵,					- 8	. 115	1 22	- 01				_ +1

		50					55					60				
5	Ser 65	Ser	His	Ser	Gln	Ser 70	Gly	Ser	Gly	Gly	Ile 75	Phe	Pro	Pro	Pro	Pro 80
	Asp	Asp	Asp	Ile	Туг 85	Asp	Gly	Ile	Glu	Glu 90	Glu	Asp	Ala	Asp	Asp 95	Gly
10	Phe	Pro	Ala	Pro 100	Pro	Lys	Gln	Leu	Asp 105	Met	Gly	Asp	Glu	Val 110	Tyr	Asp
	Asp	Val	Asp 115	Thr	Ser	Asp	Phe	Pro 120	Val	Ser	Ser	Ala	Glu 125	Met	Ser	Gln
15	Gly	Thr 130	Asn	Val	Gly	Lys	Ala 135	Lys	Thr	Glu	Glu	Lys 140	Asp	Leu	Lys	Lys
20	Leu 145	Lys	Lys	Gln	Xaa	Lys 150	Glu	Xaa	Lys	qzA	Phe 155	Arg	Lys	Lys	Phe	L ys 160
	Tyr	Asp	Gly	Glu	Ile 165	Arg	Val	Leu	Tyr	Ser 170	Thr	Lys	Val	Thr	Thr 175	Ser
25	Ile	Thr	Ser	Lys 180	Lys	Trp	Gly	Thr	Arg 185	Asp	Leu	Gln	Val	Lys 190	Pro	Gly
	Glu	Ser	Leu 195	Glu	Val	Ile	Gln	Thr 200	Thr	Asp	Asp	Thr	Lys 205	Val	Leu	Cys
30		210					215					220	Ser			
35	Asp 225	Asn	Asp	Gly	Glu	Ile 230	Tyr	Asp	Asp	Ile	Ala 235	Asp	Gly	Cys	Ile	Туг 240
		Asn	-													
40	(2)	INF		SEQU)	FOR ENCE A) L B) T	CHA ENGT	RACT H: 1	ERIS	TICS		.ds					
45			(xi)		D) I					EQ I	D NC	: 32	3:			
50	Ser 1	Met	Ser	Ala	Leu 5	Thr	Arg	Leu	Ala	Ser 10		Ala	Arg	Val	Gly 15	Gly
	Arg	Leu	Phe	Arg 20		Gly	Cys	Ala	Arg 25		Ala	Gly	Asp	Gly 30		Val
55	Arg	His	Ala 35	Gly	Gly	Gly	Val	His 40	Ile	Glu	Pro	Arg	Tyr 45		Gln	Phe
	Pro	Gln 50		Thr	Arg	Ser	Gln 55		Phe	Gln	Ser	Glu 60	Phe	Phe	Ser	Gly
60	Leu	Met	Trp	Phe	Trp	Ile	Leu	Trp	Arg	Phe	Trp	His	Asp	Ser	Glu	Glu

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65 70 75 80

Glu Leu Gly Ile Pro Pro Asp Asp Glu Asp 100 105

Applicant's or agent's tile	LOO4PCT	International application	Unassigned
reference number		1	

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorganis on page 73 . lin	,
. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
lame of depositary institution American Type Cul	lture Collection
Address of depositary institution (including postal code a	and country)
10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	
Date of deposit March 7, 1997	Accession Number 97923
C. ADDITIONAL INDICATIONS (leave blank if no	of applicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDI	ICATIONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICATIO	NS (leave blank ij noi appiicable)
The indications listed below will be submitted to the Inte Number of Deposit	emational Rureau later renegio, the general nature of the indications e.g. "Accession
	The general nature in the indicates of the second
For receiving Office use only	For International Bureau use only
	For International Bureau use only

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Applicant's or agent's tile Z004PCT	International application. Unassigned
reference number	The mational application. Ottassigney.
reterence number	
	The second secon

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorganis on page 73 . line	
3. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Cult	ture Collection
ddress of depositary institution (including postal code ar	nd country)
0801 University Boulevard Manassas. Virginia 20110-2209 United States of America	
Date of deposit May 22, 1997	Accession Number 209071
C. ADDITIONAL INDICATIONS (leave blank if no	or applicable) This information is continued on an additional sheet
DESIGNATED STATES FOR WHICH INDI	CATIONS ARE MADE (if the indications are not for all designated States
E. SEPARATE FURNISHING OF INDICATIO	
The indications listed below will be submitted to the Inter Number of Deposit')	rmational Bureau later (specify the general nature of the indications, e.g., "Access
For receiving Office use only	For International Bureau use only
This sheet was received with the international application	This sheet was received by the International Bureau on:
Authorized officer	Authorized officer

Applicant's or agent's file reference number	Z004PCT	International application	Unassigned	

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorganism referred to in the description on page 73 line N/A	
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet [17]
Name of depositary institution American Type Culture Co	ollection
Address of depositary institution tincluding postal code and coun	ury)
10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	
Date of deposit February 25, 1998	Accession Number 209641
C. ADDITIONAL INDICATIONS (leave blank if not applica-	able) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICATE	ONE ARE MARK
b. Designated States for which indicate	ONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICATIONS (lea	ive blank if not applicable)
	al Bureau later (specify the general nature of the indications, e.g., "Accession
For each in a Office	
For receiving Office use only	For International Bureau use only
This sheet was received with the international application	This sheet was received by the International Bureau on:
Authorized officer	Authorized officer
Juginia I lily	11

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5	•	1	

Applicant's or agent's file ZO reference number	04PCT		Unassigned	
	•	in resemble 6 .		

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorganism referre on page 75 . line N/A	
3. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet :
Name of depositary institution American Type Culture Coll	lection
Address of depositary institution (including postal code and country	(ش
10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	
Date of deposit July 24, 1997	Accession Number 209179
C. ADDITIONAL INDICATIONS (leave blank if not applicable)	blei This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICATION E. SEPARATE FURNISHING OF INDICATIONS (learn	ONS ARE MADE (if the indications are not for all designated States)
The indications listed below will be submitted to the Internationa Number of Deposit")	al Burcau later (specify the general nature of the indications, e.g., ".lccessio
For receiving Office use only	For International Bureau use only
This sheet was received with the international application	This sheet was received by the International Bureau on:
Authorized officer	Authorized officer

Applicant's or agent's tile	
reference number	

Z004PCT

International application Unassigned

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorganism refined page 77 . line N	erred to in the description I/A
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet !
Name of depositary institution American Type Culture C	Collection
Address of depositary institution (including postal code and con	untry)
10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	
Date of deposit March 7, 1997	Accession Number 97924
C. ADDITIONAL INDICATIONS (leave blank if not applied	cable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICATI	ONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICATIONS (lead	ave blank if not applicable) al Burcau later (specify the general nature of the indications, e.g., "Accessic
For receiving Office use only	For International Bureau use only
This sheet was received with the international application	This sheet was received by the International Bureau on
Diramia L lelia	Authorized officer

Applicant's or agent's tile reference number	Z004PCT	International application	
		The second that was the second that we see the second that the	

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorgani on page 80 . lin	
3. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Cu	lture Collection
Address of depositary institution (<i>including postal code o</i> 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	and country)
Date of deposit March 13, 1997	Accession Number 97958
C. ADDITIONAL INDICATIONS (leave blank if no	or applicable) This information is continued on an additional sheet
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorganism referr on page 80 , line N/A	
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture Col	llection
Address of depositary institution (including postal code and count 10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	(ייָרו
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B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture C	Collection
Address of depositary institution (including postal code and coll 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	uniry)
Date of deposit September 4, 1997	Accession Number 209235
C. ADDITIONAL INDICATIONS (leave blank if not appl	olicable) This information is continued on an additional sheet
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B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture	e Collection
Address of depositary institution (including postal code and of	country)
10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	
Date of deposit August 28, 1997	
Date of deposit August 28, 1997	Accession Number 209226
C. ADDITIONAL INDICATIONS (leave blank if not ap,	pplicable) This information is continued on an additional sheet
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Name of depositary institution American Type Culture	: Collection
Address of depositary institution (including postal code and code) 10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	country)
Date of deposit March 13, 1997	Accession Number 97957
C. ADDITIONAL INDICATIONS (leave blank if not app	pplicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICA	TIONS ARE MADE (if the indications are not for all designated States)
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B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American	1 Type Culture Collection
Address of depositary institution (including p 10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	osial code and country)
Date of deposit May 22, 1997	Accession Number 209073
C. ADDITIONAL INDICATIONS (lea	ve blank if not applicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WH	ICH INDICATIONS ARE MADE (if the indications are not for all designated States)
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What Is Claimed Is:

- 1. An isolated nucleic acid molecule comprising a polynucleotide having a nucleotide sequence at least 95% identical to a sequence selected from the group consisting of:
- (a) a polynucleotide fragment of SEQ ID NO:X or a polynucleotide fragment of the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X:
 - (b) a polynucleotide encoding a polypeptide fragment of SEQ ID NO:Y or a polypeptide fragment encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEO ID NO:X;
 - (c) a polynucleotide encoding a polypeptide domain of SEQ ID NO:Y or a polypeptide domain encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
 - (d) a polynucleotide encoding a polypeptide epitope of SEQ ID NO:Y or a polypeptide epitope encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
 - (e) a polynucleotide encoding a polypeptide of SEQ ID NO:Y or the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X, having biological activity;
 - (f) a polynucleotide which is a variant of SEQ ID NO:X;
 - (g) a polynucleotide which is an allelic variant of SEO ID NO:X;
 - (h) a polynucleotide which encodes a species homologue of the SEQ ID NO:Y;
 - (i) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(h), wherein said polynucleotide does not hybridize under stringent conditions to a nucleic acid molecule having a nucleotide sequence of only A residues or of only T residues.
 - 2. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding a secreted protein.
 - The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding the sequence identified as SEQ ID NO:Y or the polypeptide encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

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4. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises the entire nucleotide sequence of SEQ ID NO:X or the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

- 5. The isolated nucleic acid molecule of claim 2, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.
- 10 6. The isolated nucleic acid molecule of claim 3, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.
- 7. A recombinant vector comprising the isolated nucleic acid molecule of claim 1.
 - 8. A method of making a recombinant host cell comprising the isolated nucleic acid molecule of claim 1.
- 20 9. A recombinant host cell produced by the method of claim 8.
 - 10. The recombinant host cell of claim 9 comprising vector sequences.
- An isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence selected from the group consisting of:
 - (a) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (b) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z, having biological activity;
- 30 (c) a polypeptide domain of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (d) a polypeptide epitope of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
- (e) a secreted form of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (f) a full length protein of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;

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- (g) a variant of SEQ ID NO:Y;
- (h) an allelic variant of SEQ ID NO:Y; or
- (i) a species homologue of the SEQ ID NO:Y.
- 12. The isolated polypeptide of claim 11, wherein the secreted form or the full length protein comprises sequential amino acid deletions from either the C-terminus or the N-terminus.
 - 13. An isolated antibody that binds specifically to the isolated polypeptide of claim 11.
 - 14. A recombinant host cell that expresses the isolated polypeptide of claim 11.
 - 15. A method of making an isolated polypeptide comprising:
- (a) culturing the recombinant host cell of claim 14 under conditions such that said polypeptide is expressed; and
 - (b) recovering said polypeptide.
 - 16. The polypeptide produced by claim 15.
 - 17. A method for preventing, treating, or ameliorating a medical condition, comprising administering to a mammalian subject a therapeutically effective amount of the polypeptide of claim 11 or the polynucleotide of claim 1.
- 25 18. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
 - (a) determining the presence or absence of a mutation in the polynucleotide of claim 1; and
- (b) diagnosing a pathological condition or a susceptibility to a pathological
 condition based on the presence or absence of said mutation.
 - 19. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
 - (a) determining the presence or amount of expression of the polypeptide of claim 11 in a biological sample; and
 - (b) diagnosing a pathological condition or a susceptibility to a pathological condition based on the presence or amount of expression of the polypeptide.

- 20. A method for identifying a binding partner to the polypeptide of claim 11 comprising:
 - (a) contacting the polypeptide of claim 11 with a binding partner; and
- 5 (b) determining whether the binding partner effects an activity of the polypeptide.
 - 21. The gene corresponding to the cDNA sequence of SEQ ID NO:Y.
- 10 22. A method of identifying an activity in a biological assay, wherein the method comprises:
 - (a) expressing SEQ ID NO:X in a cell;
 - (b) isolating the supernatant;
 - (c) detecting an activity in a biological assay; and
- (d) identifying the protein in the supernatant having the activity.
 - 23. The product produced by the method of claim 22.

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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

A. The indications made below relate to the microorganism re on page 80 . line 1	eferred to in the description N/A
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture	Collection
Address of depositary institution (including postal code and co	ountry)
10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	-
Date of deposit March 13, 1997	Accession Number 97958
C. ADDITIONAL INDICATIONS (leave blank if not app.	I clicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICAT	TIONS ARE MADE (if the indications are not for all designated States)
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Name of depositary institution American Type Culture Co	ollection
Address of depositary institution (including postal code and coun	π ό .)
10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	
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B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture Collection	on
Address of depositary institution (including postal code and country) 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	
Date of deposit September 4, 1997 Ac	ecession Number 209235
C. ADDITIONAL INDICATIONS (leave blank if not applicable)	This information is continued on an additional sheet
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B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture Co	ollection
Address of depositary institution (including postal code and cour	ntry)
10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	
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Address of depositary institution (including postal code and coll 10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	ountry)
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C. ADDITIONAL INDICATIONS (leave blank if not appl	dicable) This information is continued on an additional sheet*
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Address of depositary institution (<i>including postal code and c</i> 10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	ountry)
Date of deposit May 22, 1997	Accession Number 209073
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What Is Claimed Is:

- 1. An isolated nucleic acid molecule comprising a polynucleotide having a nucleotide sequence at least 95% identical to a sequence selected from the group consisting of:
- (a) a polynucleotide fragment of SEQ ID NO:X or a polynucleotide fragment of the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (b) a polynucleotide encoding a polypeptide fragment of SEQ ID NO:Y or a polypeptide fragment encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (c) a polynucleotide encoding a polypeptide domain of SEQ ID NO:Y or a polypeptide domain encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (d) a polynucleotide encoding a polypeptide epitope of SEQ ID NO:Y or a polypeptide epitope encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (e) a polynucleotide encoding a polypeptide of SEQ ID NO:Y or the cDNA sequence included in ATCC Deposit No:Z. which is hybridizable to SEQ ID NO:X, having biological activity;
 - (f) a polynucleotide which is a variant of SEO ID NO:X;
 - (g) a polynucleotide which is an allelic variant of SEO ID NO:X;
 - (h) a polynucleotide which encodes a species homologue of the SEQ ID NO:Y;
- (i) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(h), wherein said polynucleotide does not hybridize under stringent conditions to a nucleic acid molecule having a nucleotide sequence of only A residues or of only T residues.
 - 2. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding a secreted protein.
- 3. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding the sequence identified as SEQ ID NO:Y or the polypeptide encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

4. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises the entire nucleotide sequence of SEQ ID NO:X or the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

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5. The isolated nucleic acid molecule of claim 2, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.

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- 6. The isolated nucleic acid molecule of claim 3, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.
- 7. A recombinant vector comprising the isolated nucleic acid molecule of claim 1.
 - 8. A method of making a recombinant host cell comprising the isolated nucleic acid molecule of claim 1.

- 9. A recombinant host cell produced by the method of claim 8.
- 10. The recombinant host cell of claim 9 comprising vector sequences.
- 11. An isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence selected from the group consisting of:
 - (a) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (b) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z, having biological activity;
- 30 (c) a polypeptide domain of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (d) a polypeptide epitope of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
- (e) a secreted form of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (f) a full length protein of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;

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- (g) a variant of SEQ ID NO:Y;
- (h) an allelic variant of SEQ ID NO:Y; or
- (i) a species homologue of the SEQ ID NO:Y.
- 12. The isolated polypeptide of claim 11, wherein the secreted form or the full length protein comprises sequential amino acid deletions from either the C-terminus or the N-terminus.
 - 13. An isolated antibody that binds specifically to the isolated polypeptide of claim 11.
 - 14. A recombinant host cell that expresses the isolated polypeptide of claim 11.
 - 15. A method of making an isolated polypeptide comprising:
 - (a) culturing the recombinant host cell of claim 14 under conditions such that said polypeptide is expressed; and
 - (b) recovering said polypeptide.
 - 16. The polypeptide produced by claim 15.
 - 17. A method for preventing, treating, or ameliorating a medical condition, comprising administering to a mammalian subject a therapeutically effective amount of the polypeptide of claim 11 or the polynucleotide of claim 1.
- 25 18. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
 - (a) determining the presence or absence of a mutation in the polynucleotide of claim 1; and
- (b) diagnosing a pathological condition or a susceptibility to a pathologicalcondition based on the presence or absence of said mutation.
 - 19. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
- (a) determining the presence or amount of expression of the polypeptide of claim 11 in a biological sample; and
 - (b) diagnosing a pathological condition or a susceptibility to a pathological condition based on the presence or amount of expression of the polypeptide.

- 20. A method for identifying a binding partner to the polypeptide of claim 11 comprising:
 - (a) contacting the polypeptide of claim 11 with a binding partner; and
- 5 (b) determining whether the binding partner effects an activity of the polypeptide.
 - 21. The gene corresponding to the cDNA sequence of SEQ ID NO:Y.
- 10 22. A method of identifying an activity in a biological assay, wherein the method comprises:
 - (a) expressing SEQ ID NO:X in a cell;
 - (b) isolating the supernatant;
 - (c) detecting an activity in a biological assay; and
- (d) identifying the protein in the supernatant having the activity.
 - 23. The product produced by the method of claim 22.